

Medical Dictionary: Hepatocellular carcinoma (fibrolamellar variant)

A B C D E F G H I J K L M N O P Q R S T U V W X Y Z #

Hepatocellular carcinoma (fibrolamellar variant): Rare Disease

Office of Rare Diseases (ORD) of the National Institutes of Health (NIH)

Hepatocellular carcinoma (fibrolamellar variant) is listed as a "rare disease" by the Office of Rare Diseases of the National Institutes of Health (NIH). This means that Hepatocellular carcinoma (fibrolamellar variant), or a subtype of (fibrolamellar variant), affects less than 200,000 people in the US population.

Source - [National Institutes of Health \(NIH\)](#)

Terms associated with Hepatocellular carcinoma (fibrolamellar variant):

Terms that may be interchangeable with Hepatocellular carcinoma (fibrolamellar variant)

- [Fibrolamellar hepatocellular carcinoma](#)
- [Fibrolamellar variant of hepatocellular carcinoma](#)

Source - NIH

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Last updated: 15 November, 2006

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2006 ACS on STN

RN 114977-28-5 REGISTRY

ED Entered STN: 25 Jun 1988

CN Benzenepropanoic acid, β -[[(1,1-dimethylethoxy) carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 7,11-Methano-1H-cyclodeca[3,4]benz[1,2-b]oxete, benzenepropanoic acid deriv.

CN Benzenepropanoic acid, β -[[(1,1-dimethylethoxy) carbonyl]amino]- α -hydroxy-, 12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, [2aR-[2a α ,4 β ,4a β ,6 β ,9 α (α R*, β S*)],11 α ,12 α ,12a α ,12b α]-

OTHER NAMES:

CN Docetaxel

CN Docetaxol

CN N-Debenzoyl-N-tert-butoxycarbonyl-10-deacetyltaxol

CN RP 56976

CN Taxotere

FS STEREOSEARCH

DR 216252-50-5

MF C43 H53 N O14

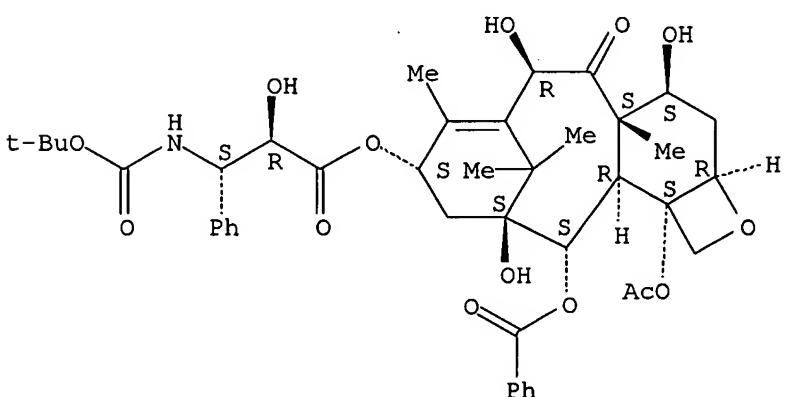
CI COM

SR CA

LC STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOSIS, BIOTECHNO, CA, CAPLUS, CASREACT; CBNB, CHEMCATS, CHEMINFORMRX, CIN, CSCHEM, DDFU, DRUGU, EMBASE, HSDB*, IMSCOSEARCH, IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK*, MSDS-OHS, PATDPASPC, PHAR, PIRA, PROMT, PROUSDDR, PS, RTECS*, SYNTHLINE, TOXCENTER, USAN, USPAT2, USPATFULL

(*File contains numerically searchable property data)

Absolute stereochemistry.



PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

3431 REFERENCES IN FILE CA (1907 TO DATE)

132 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

3480 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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NEWS 10 SEP 25 CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine
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NEWS 21 NOV 13 CA/CAplus pre-1967 chemical substance index entries enhanced with preparation role
NEWS 22 NOV 20 CAS Registry Number crossover limit increased to 300,000 in additional databases
NEWS 23 NOV 20 CA/CAplus to MARPAT accession number crossover limit increased to 50,000
NEWS 24 NOV 20 CA/CAplus patent kind codes will be updated
NEWS 25 DEC 01 CAS REGISTRY updated with new ambiguity codes

NEWS EXPRESS NOVEMBER 10 CURRENT WINDOWS VERSION IS V8.01c, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 25 SEPTEMBER 2006.

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=> focus
PROCESSING COMPLETED FOR L9
L10 420 FOCUS L9 1-

=> s l10 and pd<=2000
2 FILES SEARCHED...
L11 55 L10 AND PD<=2000

=> focus
PROCESSING COMPLETED FOR L11
L12 55 FOCUS L11 1-

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L12 ANSWER 1 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2000:650912 CAPLUS
DOCUMENT NUMBER: 134:141449
TITLE: Comparison of 2-methoxyestradiol-induced, docetaxel-induced, and paclitaxel-induced apoptosis in hepatoma cells and its correlation with reactive oxygen species
AUTHOR(S): Lin, Heng-Liang; Liu, Tsung-Yun; Chau, Gar-Yang; Lui, Wing-Yiu; Chi, Chin-Wen
CORPORATE SOURCE: Institute of Pharmacology, National Yang-Ming University, Taipei, Taiwan
SOURCE: Cancer (New York) (2000), 89(5), 983-994
CODEN: CANCAR; ISSN: 0008-543X
PUBLISHER: John Wiley & Sons, Inc.
DOCUMENT TYPE: Journal
LANGUAGE: English
AB Previously, the authors observed that paclitaxel treatment of hepatoma cells resulted in differential cytotoxicity. Whether other antimicrotubule agents (docetaxel and 2-methoxyestradiol) are more effective than paclitaxel is not clear. Moreover, whether the modulation of reactive oxygen species (ROS) is involved in the drug-induced growth inhibition of hepatoma cells is not known. The authors examined the effects of 2-methoxyestradiol, paclitaxel, and docetaxel on HepG2, Hep3B, HA22T/VGH, and Hepal-6 hepatoma cell lines. The parameters examined included cell viability, cell membrane permeability, cell cycle distribution, DNA fragmentation, and ROS generation. Docetaxel and paclitaxel inhibited the growth of hepatoma cells at submicromolar concns., whereas that of 2-methoxyestradiol was within a micromolar range. This drug-induced growth inhibition was cell cycle dependent. 2-Methoxyestradiol-treated (10-50 µM) cells resulted in G2/M block prior to apoptosis. High dose (0.1 µM) docetaxel- and paclitaxel-treated cells resulted in a G2/M arrest followed by generation of polyploidy or apoptosis; however, low dose (0.01 µM) treatment induced apoptosis without G2/M arrest. The low dose effect was more significant in docetaxel-treated cells than in paclitaxel-treated cells. Although these antimicrotubule agents increased the formation of ROS, antioxidant treatment did not block drug-induced cell cycle and growth inhibition effects. The current results suggest that the growth inhibition of hepatoma cells induced by 2-methoxyestradiol, paclitaxel, and docetaxel was mediated through G2/M-phase arrest, caspase activation, and DNA fragmentation. The drug-induced apoptosis was independent of ROS formation. Docetaxel was more effective than paclitaxel in killing hepatoma

cells. The potential of using 2-methoxyestradiol and docetaxel for the treatment of patients with hepatoma is worthy of further study.

IT 114977-28-5, Docetaxel

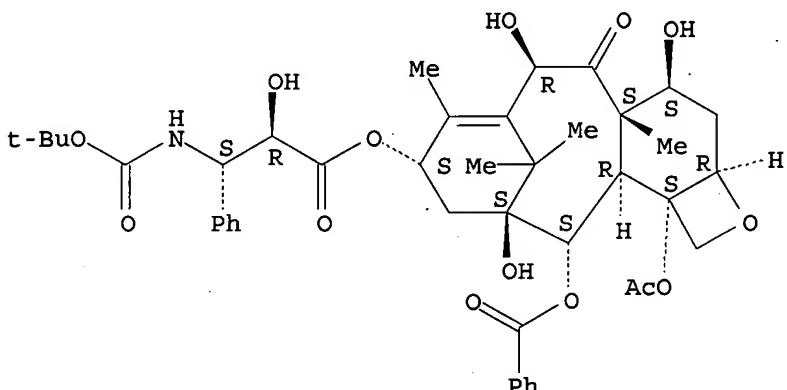
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(2-methoxyestradiol-, docetaxel-, and paclitaxel-induced apoptosis in hepatoma cells)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S) - (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 58 THERE ARE 58 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 2 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:126569 CAPLUS

DOCUMENT NUMBER: 132:175461

TITLE: Factors predicting for efficacy and safety of docetaxel in a compassionate-use cohort of 825 heavily pretreated advanced breast cancer patients

AUTHOR(S): Alexandre, J.; Bleuzen, P.; Bonneterre, J.; Sutherland, W.; Misset, J. L.; Guastalla, J.-P.; Viens, P.; Faivre, S.; Chahine, A.; Spielman, M.; Bensmaïne, A.; Marty, M.; Mahjoubi, M.; Cvitkovic, E.

CORPORATE SOURCE: Paul Brousse Hospital and Institut Gustave Roussy, Villejuif, 94804, Fr.

SOURCE: Journal of Clinical Oncology (2000), 18(3), 562-573

PUBLISHER: CODEN: JCONDN; ISSN: 0732-183X

DOCUMENT TYPE: Lippincott Williams & Wilkins

LANGUAGE: English

AB Purpose: To identify predictive factors for efficacy and safety in advanced breast cancer (ABC) patients treated in the French compassionate-use docetaxel program. Patients and Methods: A total of 825 ABC patients treated with docetaxel (100 mg/m² every 3 wk) were source-reviewed and analyzed for prognostic factors associated with overall response rate (ORR), time to treatment failure (TTF), overall survival (OS), febrile neutropenia, mucositis, and severe fluid retention syndrome by univariate and multivariate anal. Results: The ORR was 22.9% (95% confidence interval, 20.2% to 26.2%). The median TTF and

OS were 4.0 and 9.8 mo, resp. By multivariate anal., secondary anthracycline-resistant disease was significantly associated ($P < .05$) with lower ORR and shorter TTF and OS, whereas anthracycline-refractory disease was associated with shorter OS. Poor performance status was associated with lower ORR, shorter TTF, and shorter OS. Liver dysfunction (transaminase levels > 1.5 times the upper limit of normal [ULN] and alkaline phosphatase [AP] level $>$ three times ULN) and time since first relapse less than 24 mo were associated with shorter TTF and OS. Other significant correlations included the following: elevated CA 15-3 serum level with lower ORR; more than two involved sites, and minor transaminase and AP level abnormalities with shorter OS; and no previous chemotherapy for ABC with shorter TTF. According to multivariate anal., ORR, TTF, and OS were not decreased in patients with liver metastases but without liver dysfunction. Conclusion: Docetaxel activity was maintained in heavily pretreated ABC patients and in those with liver metastasis; docetaxel must be used cautiously, however, in patients with liver dysfunction in whom high morbidity risk necessitates strict adherence to dose-adaptation guidelines.

IT

114977-28-5, Docetaxel

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(factors predicting for efficacy and safety of docetaxel in
pretreated humans with advanced breast cancer)

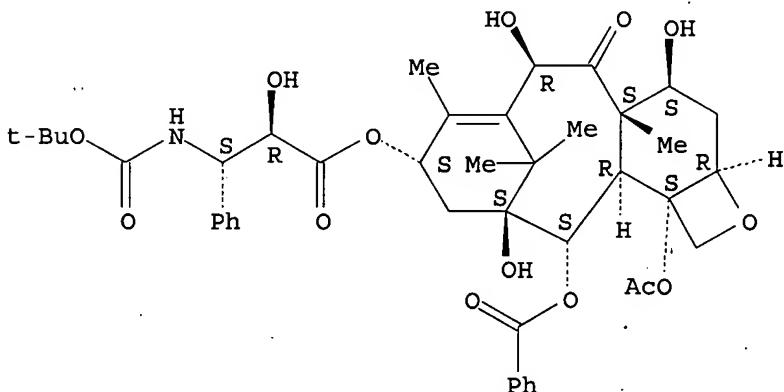
RN

114977-28-5 CAPLUS

CN

Benzene propanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT:

40

THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 3 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:60933 CAPLUS

DOCUMENT NUMBER: 132:102535

TITLE: A Phase I study of gemcitabine and docetaxel in patients with metastatic solid tumors

AUTHOR(S): Ryan, David P.; Lynch, Thomas J.; Grossbard, Michael L.; Seiden, Michael V.; Fuchs, Charles S.; Grenon, Nina; Baccala, Paul; Berg, Deborah; Finkelstein, Dianne; Mayer, Robert J.; Clark, Jeffrey W.

CORPORATE SOURCE: Gastrointestinal Cancer Clinic, Dana-Farber/Partners CancerCare, Boston, MA, 02114, USA

SOURCE: Cancer (New York) (2000), 88(1), 180-185

CODEN: CANCAR; ISSN: 0008-543X

PUBLISHER: John Wiley & Sons, Inc.
 DOCUMENT TYPE: Journal
 LANGUAGE: English
 AB A Phase I study was initiated to determine the maximum tolerated dose of weekly gemcitabine combined with monthly, fixed-dose docetaxel. Patients with metastatic solid tumors were treated with docetaxel, 60 mg/m², on Day 1 every 28 days. Gemcitabine was administered on Days 1, 8, and 15 and underwent dose adjustment in cohorts of 3-6 patients. At the maximum tolerated dose, 11 addnl. patients were enrolled. Twenty-six patients received 85 cycles of therapy. At the first dose level, the planned gemcitabine dose on Days 1, 8, and 15 was 800 mg/m². Two of the 6 patients treated at this dose level experienced dose-limiting toxicities (DLTs) requiring the reduction of gemcitabine to 600 mg/m² per dose and the administration of ciprofloxacin, 500 mg orally twice daily, on Days 8-18. At the second dose level the first 3 patients experienced no DLTs and the dose of gemcitabine was increased to 700 mg/m². Two of the 6 patients treated at the 700 mg/m² dose level experienced DLTs. Eleven addnl. patients were enrolled at the recommended Phase II dose of gemcitabine (600 mg/m²). At this dose level, Grade 3/4 (according the National Cancer Institute's common toxicity criteria) neutropenia and thrombocytopenia occurred in 12.5% and 2.1% of cycles, resp. Grade 3 and 4 nonhematol. toxicities were uncommon. Three of seven evaluable patients with pancreatic carcinoma had evidence of significant antineoplastic activity (three partial responses). In addition, two complete responses (one patient with gastric carcinoma and one patient with ovarian carcinoma) and one partial response (patient with hepatocellular carcinoma) were noted in patients with other solid tumors. The regimen comprised of docetaxel, 60 mg/m², on Day 1 and gemcitabine, 600 mg/m², on Days 1, 8, and 15 with ciprofloxacin on Days 8-18 every 28 days is safe, well tolerated, and active.

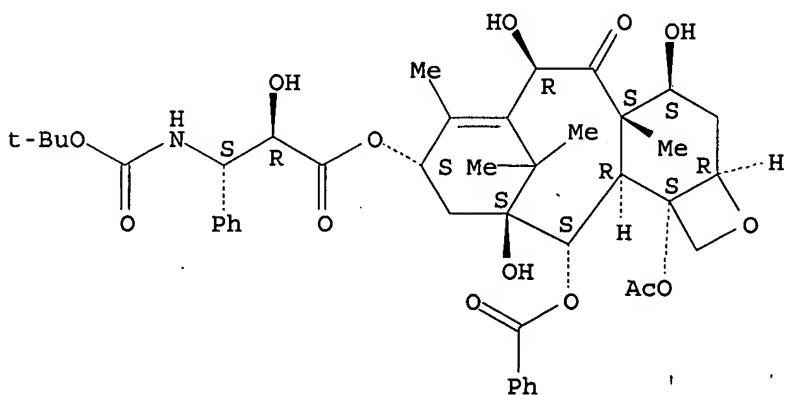
IT 114977-28-5, Docetaxel

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (gemcitabine and docetaxel in human patients with metastatic solid tumors)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a;5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT:

26

THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 4 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1999:641123 CAPLUS

DOCUMENT NUMBER: 132:117128

TITLE: Efficacy and safety of docetaxel (Taxotere)

in heavily pretreated advanced breast cancer patients; the French compassionate use program experience

AUTHOR(S): Bonneterre, J.; Spielman, M.; Guastalla, J. -P.;

Marty, M.; Viens, P.; Chollet, P.; Roche, H.;

Fumoleau, P.; Mauriac, L.; Bourgeois, H.; Namer, M.;

Bergerat, J. P.; Misset, J. -L.; Trandafir, L.;

Mahjoubi, M.

CORPORATE SOURCE: Centre Oscar Lambret, Lille, 59020, Fr.

SOURCE: European Journal of Cancer (1999), 35(10),

1431-1439

CODEN: EJCAEL; ISSN: 0959-8049

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal

LANGUAGE: English

AB The aim was to assess retrospectively doceaxel safety and efficacy in advanced breast cancer patients in a French compassionate use program. Patients had received >1 prior chemotherapy regimen for advanced disease, were either anthracycline-resistant (that is progressed within 6 mo after anthracycline-based chemotherapy) or had received the maximum cumulative dose. The recommended docetaxel dose was 100 mg/m²/cycle (75 mg/m²) prior palliative chemotherapy lines. The most frequent severe toxicity, febrile neutropenia (reported in 223/870 (25.6%) patients evaluable for safety), caused 10 deaths, 6 of these being patients with severe liver impairment before inclusion. Fluid retention syndrome and other common non-Hematol. toxicities were well tolerated. 3.1% (28/889) of all patients and 11.4% of those with liver dysfunction, died from treatment-related causes. The overall response rate in 825 assessable patients was 22.9% (95% confidence interval (CI): 20.2-26.2%). Median time to treatment failure was 4 mo (95% CI: 3.6-4.3) and median survival was 9.8 mo (95% CI: 8.8-10.7). This report on the largest series of unselected advanced breast cancer patients treated with docetaxel supports previous phase II studies, confirming docetaxel's utility in patients relapsing after failing anthracycline-containing palliative chemotherapy.

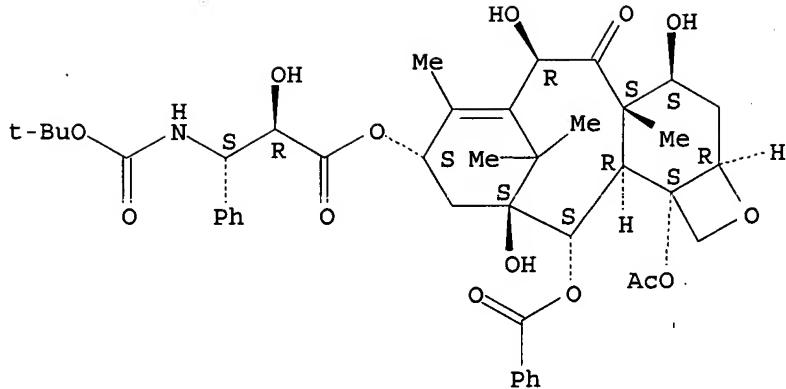
IT 114977-28-5, Docetaxel

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(efficacy and safety of docetaxel in heavily pretreated advanced breast cancer patients)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 5 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:790283 CAPLUS

DOCUMENT NUMBER: 133:344606

TITLE: Combined pharmaceuticals comprising anthracycline derivatives

INVENTOR(S): Geroni, Maria Cristina; Ripamonti, Marina; Caruso, Michele; Suarato, Antonino

PATENT ASSIGNEE(S): Pharmacia & Upjohn S.p.A., Italy

SOURCE: PCT Int. Appl., 23 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|--------------|
| WO 2000066093 | A2 | 20001109 | WO 2000-EP2923 | 20000404 <-- |
| WO 2000066093 | A3 | 20010125 | | |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| EP 1173187 | A2 | 20020123 | EP 2000-925158 | 20000404 |
| EP 1173187 | B1 | 20030806 | | |
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| JP 2002543112 | T2 | 20021217 | JP 2000-614978 | 20000404 |
| EP 1323423 | A1 | 20030702 | EP 2003-75776 | 20000404 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL | | | | |
| AT 246507 | E | 20030815 | AT 2000-925158 | 20000404 |
| PT 1173187 | T | 20031031 | PT 2000-925158 | 20000404 |
| ES 2204572 | T3 | 20040501 | ES 2000-925158 | 20000404 |
| CN 1507869 | A | 20040630 | CN 2003-10114911 | 20000404 |
| CN 1535688 | A | 20041013 | CN 2003-10114909 | 20000404 |
| CN 1853645 | A | 20061101 | CN 2006-10059701 | 20000404 |
| CN 1853642 | A | 20061101 | CN 2006-10059703 | 20000404 |
| TW 222863 | B1 | 20041101 | TW 2000-89106805 | 20000412 |
| US 6537990 | B1 | 20030325 | US 2001-926392 | 20011025 |
| HK 1045462 | A1 | 20060428 | HK 2002-107029 | 20020926 |

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|------------------------|-------------|------------------|-------------|
| US 2003087839 | A1 20030508 | US 2002-284144 | 20021031 |
| US 6586428 | B2 20030701 | | |
| PRIORITY APPLN. INFO.: | | | |
| | | GB 1999-9925 | A 19990429 |
| | | CN 2000-806897 | A 20000404 |
| | | CN 2003-10114909 | A3 20000404 |
| | | CN 2003-10114911 | A3 20000404 |
| | | EP 2000-925158 | A3 20000404 |
| | | WO 2000-EP2923 | W 20000404 |
| | | US 2001-926392 | A1 20011025 |

AB The present invention relates to combined pharmaceuticals comprising a morpholinylanthracycline administered in combination anticancer agents chosen from an alkylating agent, an antimetabolite, a topoisomerase II inhibitor, a topoisomerase I inhibitor, an antimitotic drug and a platinum derivative, which are useful in anticancer therapy, particularly in the treatment of a primary or metastatic liver cancer. At doses 5.9 and 7.7 mg/kg cis-platin administered alone and 0.05 mg/kg PNU-152243 (morpholinylanthracycline) administered alone, the increase in lifespan values was 33, 33 and 50%. Sequential administration of these drugs showed a therapeutic advantage of the combination in comparison with each drug administered alone.

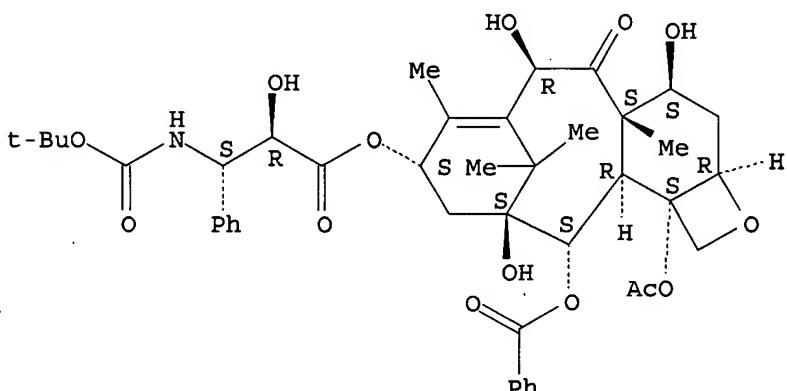
IT 114977-28-5, Docetaxel

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(combined pharmaceuticals comprising anthracycline derivs.)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 6 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:456927 CAPLUS

DOCUMENT NUMBER: 133:84243

TITLE: Method of using a cyclooxygenase-2 inhibitor and one or more antineoplastic agents as a combination therapy in the treatment of neoplasia

INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.

PATENT ASSIGNEE(S): G.D. Searle and Co., USA

SOURCE: PCT Int. Appl., 236 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 21

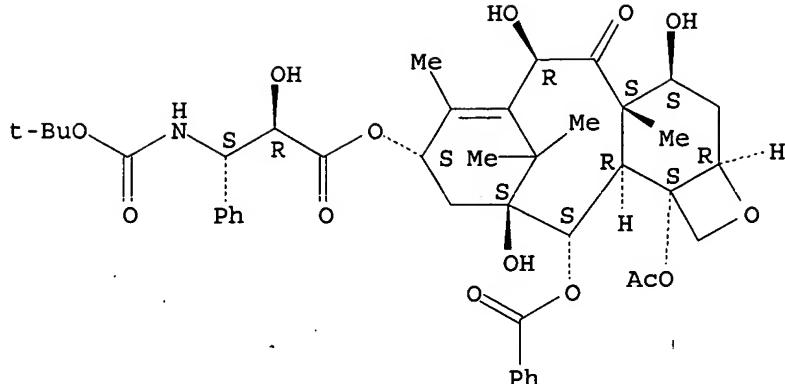
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-------------------|--------------|
| WO 2000038730 | A2 | 20000706 | WO 1999-US30693 | 19991222 <-- |
| WO 2000038730 | A3 | 20001102 | | |
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CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
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| AU 2000023805 | A5 | 20000731 | AU 2000-23805 | 19991222 <-- |
| AU 783992 | B2 | 20060112 | | |
| EP 1140192 | A2 | 20011010 | EP 1999-967543 | 19991222 |
| EP 1140192 | B1 | 20060405 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, CY | | | | |
| TR 200102499 | T2 | 20011221 | TR 2001-200102499 | 19991222 |
| BR 9916518 | A | 20020129 | BR 1999-16518 | 19991222 |
| HU 200104814 | A2 | 20020429 | HU 2001-4814 | 19991222 |
| JP 2002533416 | T2 | 20021008 | JP 2000-590681 | 19991222 |
| EP 1522313 | A1 | 20050413 | EP 2004-26577 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI, RO, CY | | | | |
| AT 322290 | E | 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A | 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A | 20020107 | ZA 2001-5120 | 20010621 |
| NO 2001003155 | A | 20010822 | NO 2001-3155 | 20010622 |
| US 2003119895 | A1 | 20030626 | US 2002-150546 | 20020516 |
| US 2003203956 | A1 | 20031030 | US 2002-212523 | 20020805 |
| AU 2004210578 | A1 | 20041007 | AU 2004-210578 | 20040910 |
| US 2005037090 | A1 | 20050217 | US 2004-945422 | 20040920 |
| US 1998-113786P | P | 19981223 | | |
| US 1999-385214 | A | 19990827 | | |
| AU 2000-25936 | A3 | 19991222 | | |
| EP 1999-968939 | A3 | 19991222 | | |
| WO 1999-US30693 | W | 19991222 | | |
| US 2001-857873 | A2 | 20011005 | | |

PRIORITY APPLN. INFO.:

- AB Methods are provided to treat or prevent neoplasia disorders in a mammal using a combination of a cyclooxygenase-2 inhibitor and an antineoplastic agent.
- IT 114977-28-5, Docetaxel
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(cyclooxygenase-2 inhibitor-antineoplastic agent combination for neoplasia treatment)
- RN 114977-28-5 CAPLUS
- CN Benzenepropanoic acid, β -[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 7 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:456915 CAPLUS

DOCUMENT NUMBER: 133:84242

TITLE: Method of using a matrix metalloproteinase inhibitor and one or more antineoplastic agents as a combination therapy in the treatment of neoplasia

INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.

PATENT ASSIGNEE(S): G.D. Searle and Co., USA

SOURCE: PCT Int. Appl., 277 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 21

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-------------------|-------------|
| WO 2000038718 | A2 | 20000706 | WO 1999-US30699 | 19991222 <- |
| WO 2000038718 | A3 | 20001109 | | |
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RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2356929 | AA | 20000706 | CA 1999-2356929 | 19991222 <- |
| AU 2000027135 | A5 | 20000731 | AU 2000-27135 | 19991222 <- |
| EP 1140182 | A2 | 20011010 | EP 1999-968941 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| TR 200102499 | T2 | 20011221 | TR 2001-200102499 | 19991222 |
| JP 2002533406 | T2 | 20021008 | JP 2000-590669 | 19991222 |
| EP 1522313 | A1 | 20050413 | EP 2004-26577 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, RO, CY | | | | |
| AT 322290 | E | 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A | 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A | 20020107 | ZA 2001-5120 | 20010621 |
| US 6858598 | B1 | 20050222 | US 2001-857995 | 20011005 |
| AU 2004210578 | A1 | 20041007 | AU 2004-210578 | 20040910 |
| US 2005058725 | A1 | 20050317 | US 2004-945002 | 20040920 |
| US 6916800 | B2 | 20050712 | | |
| PRIORITY APPLN. INFO.: | | | US 1998-113786P | P 19981223 |

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| US 1999-385214 | A 19990827 |
| AU 2000-25936 | A3 19991222 |
| EP 1999-968939 | A3 19991222 |
| WO 1999-US30699 | W 19991222 |
| US 2001-857995 | A1 20011005 |

AB Methods are provided for the prevention and treatment of neoplasia disorders in a mammal using a combination of a matrix metalloproteinase inhibitor and an antineoplastic agent.

IT 114977-28-5, Docetaxel

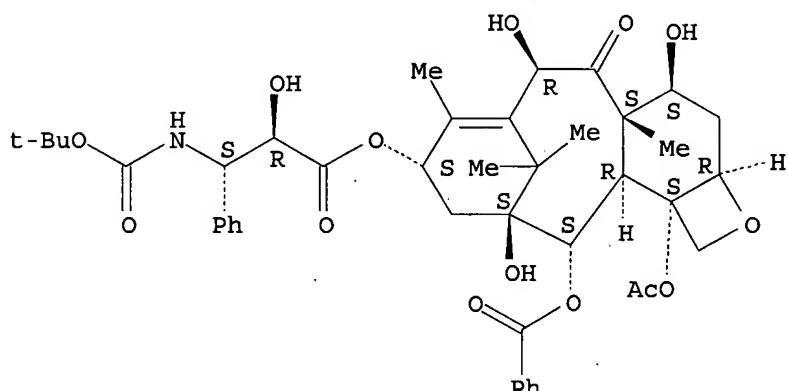
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(matrix metalloproteinase inhibitor and antineoplastic agent as combination therapy in neoplasia treatment)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 8 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:456866 CAPLUS

DOCUMENT NUMBER: 133:84239

TITLE: Method of using an integrin antagonist and one or more antineoplastic agents as a combination therapy in the treatment of neoplasia

INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.

PATENT ASSIGNEE(S): G.D. Searle and Co., USA

SOURCE: PCT Int. Appl., 220 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 21

PATENT INFORMATION:

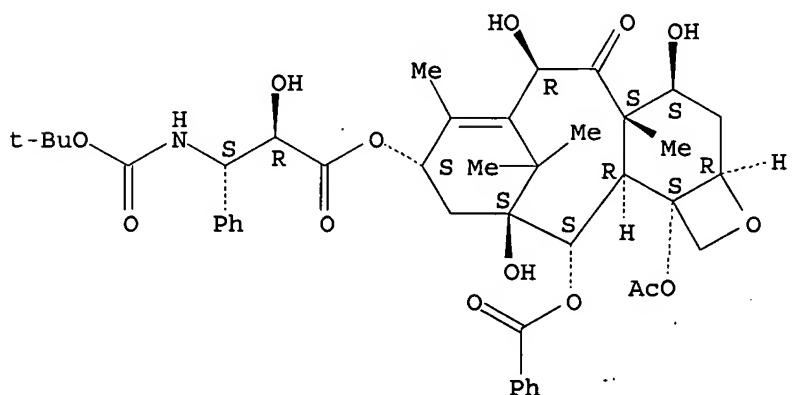
| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
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| WO 2000038665 | A2 | 20000706 | WO 1999-US30670 | 19991222 <- |
| WO 2000038665 | A3 | 20001116 | | |

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IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
 MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
 SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW
 RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
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 CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
 CA 2356462 AA 20000706 CA 1999-2356462 19991222 <--
 AU 2000025926 A5 20000731 AU 2000-25926 19991222 <--
 EP 1140193 A2 20011010 EP 1999-968529 19991222
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO
 TR 200102499 T2 20011221 TR 2001-200102499 19991222
 JP 2002533387 T2 20021008 JP 2000-590619 19991222
 EP 1522313 A1 20050413 EP 2004-26577 19991222
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, FI, RO, CY
 AT 322290 E 20060415 AT 1999-967543 19991222
 ZA 2001005055 A 20020920 ZA 2001-5055 20010620
 ZA 2001005120 A 20020107 ZA 2001-5120 20010621
 US 6833373 B1 20041221 US 2001-857994 20011005
 US 2004234624 A1 20041125 US 2004-865414 20040610
 AU 2004210578 A1 20041007 AU 2004-210578 20040910
 PRIORITY APPLN. INFO.: US 1998-113786P P 19981223
 US 1999-385214 A 19990827
 AU 2000-25936 A3 19991222
 EP 1999-968939 A3 19991222
 WO 1999-US30670 W 19991222
 US 2001-857994 A1 20011005

- AB The present invention provides methods to treat or prevent neoplasia disorders in a mammal using a combination of an integrin antagonist and an antineoplastic agent.
 IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (integrin antagonist-antineoplastic agent combination for neoplasia treatment)
 RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

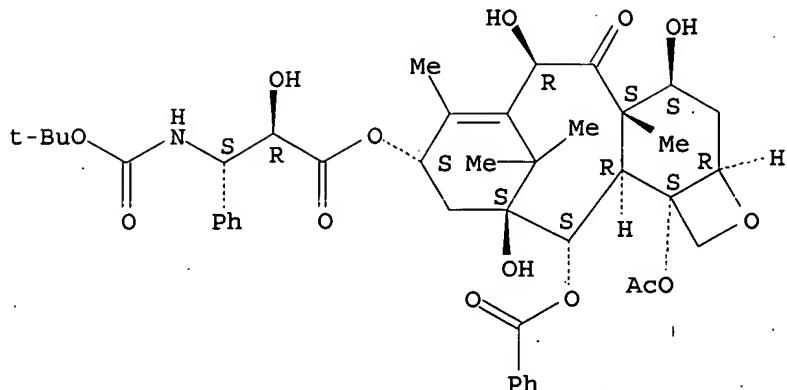


L12 ANSWER 9 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2000:368032 CAPLUS
 DOCUMENT NUMBER: 133:26843
 TITLE: Methods and compositions for diagnosis and treatment
 of cancer based on the transcription factor ets2
 INVENTOR(S): Papas, Takis S.; Watson, Dennis K.
 PATENT ASSIGNEE(S): Musc Foundation for Research Development, USA; Papas,
 Tula Christy
 SOURCE: PCT Int. Appl., 101 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 2000030590 | A2 | 20000602 | WO 1999-US27805 | 19991123 <-- |
| WO 2000030590 | A3 | 20000817 | | |
| W: AU, CA, JP, US | | | | |
| RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
PT, SE | | | | |
| CA 2351627 | AA | 20000602 | CA 1999-2351627 | 19991123 <-- |
| AU 2000024740 | A5 | 20000613 | AU 2000-24740 | 19991123 <-- |
| EP 1133575 | A2 | 20010919 | EP 1999-968046 | 19991123 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI | | | | |
| JP 2002530102 | T2 | 20020917 | JP 2000-583475 | 19991123 |
| US 2002081601 | A1 | 20020627 | US 2001-841963 | 20010425 |
| US 2004047845 | A1 | 20040311 | US 2001-841960 | 20010425 |
| PRIORITY APPLN. INFO.: | | | US 1998-109850P | P 19981125 |
| | | | WO 1999-US27805 | W 19991123 |

- AB The present invention relates to methods for treating and preventing cancer by modifying the expression of ets2 gene expression or the activity of the gene product. The invention also relates to sensitizing cancer cells to chemotherapeutic or radiotherapeutic agents. Ets2 gene expression and/or activity of the gene product can be modulated using antisense ets2 nucleic acids and/or modified ets2 proteins. The present invention also provides pharmaceutical compns. which comprise antisense ets2 nucleic acid, and nucleic acid that encode modified ets2 proteins and/or modified ets2 proteins.
- IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (diagnosis and treatment of cancer based on the transcription factor ets2)
- RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 10 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:456950 CAPLUS

DOCUMENT NUMBER: 133:84244

TITLE: Method of using a cyclooxygenase-2 inhibitor and an integrin antagonist as a combination therapy in the treatment of neoplasia

INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.

PATENT ASSIGNEE(S): G.D. Searle and Co., USA

SOURCE: PCT Int. Appl., 348 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

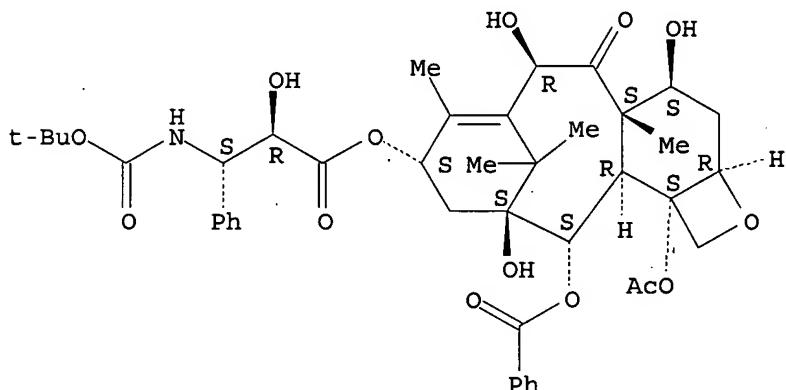
FAMILY ACC. NUM. COUNT: 21

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-------------------|-------------|
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| WO 2000038786 | A3 | 20010308 | | |
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| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2356302 | AA | 20000706 | CA 1999-2356302 | 19991222 <- |
| AU 2000022104 | A5 | 20000731 | AU 2000-22104 | 19991222 <- |
| EP 1140179 | A2 | 20011010 | EP 1999-966594 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| TR 200102499 | T2 | 20011221 | TR 2001-200102499 | 19991222 |
| JP 2002533422 | T2 | 20021008 | JP 2000-590734 | 19991222 |
| EP 1522313 | A1 | 20050413 | EP 2004-26577 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, RO, CY | | | | |
| AT 322290 | E | 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A | 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A | 20020107 | ZA 2001-5120 | 20010621 |
| AU 2004210578 | A1 | 20041007 | AU 2004-210578 | 20040910 |
| PRIORITY APPLN. INFO.: | | | US 1998-113786P | P 19981223 |
| | | | US 1999-385214 | A 19990827 |
| | | | AU 2000-25936 | A3 19991222 |
| | | | EP 1999-968939 | A3 19991222 |

- AB Methods are provided to treat or prevent neoplasia disorders in a mammal using a combination of a cyclooxygenase-2 inhibitor, an integrin antagonist and an antineoplastic agent.
- IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (cyclooxygenase-2 inhibitor and integrin antagonist in combination for neoplasia treatment)
- RN 114977-28-5 CAPLUS
- CN Benzenepropanoic acid, β -[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S).- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 11 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:456916 CAPLUS

DOCUMENT NUMBER: 133:68929

TITLE: Use of a matrix metalloproteinase inhibitor and an integrin antagonist in the treatment of neoplasia

INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.

PATENT ASSIGNEE(S): G.D. Searle and Co., USA

SOURCE: PCT Int. Appl., 358 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 21

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|-------------|
| WO 2000038719 | A1 | 20000706 | WO 1999-US30700 | 19991222 <- |
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| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |

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| AU 2000027136 | A5 20000731 | AU 2000-27136 | 19991222 <-- |
| EP 1140183 | A1 20011010 | EP 1999-968942 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO | | | |
| TR 200102499 | T2 20011221 | TR 2001-200102499 | 19991222 |
| JP 2002533407 | T2 20021008 | JP 2000-590670 | 19991222 |
| EP 1522313 | A1 20050413 | EP 2004-26577 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI, RO, CY | | | |
| AT 322290 | E 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A 20020107 | ZA 2001-5120 | 20010621 |
| AU 2004210578 | A1 20041007 | AU 2004-210578 | 20040910 |
| PRIORITY APPLN. INFO.: | | | |
| | | US 1998-113786P | P 19981223 |
| | | US 1999-385214 | A 19990827 |
| | | AU 2000-25936 | A3 19991222 |
| | | EP 1999-968939 | A3 19991222 |
| | | WO 1999-US30700 | W 19991222 |

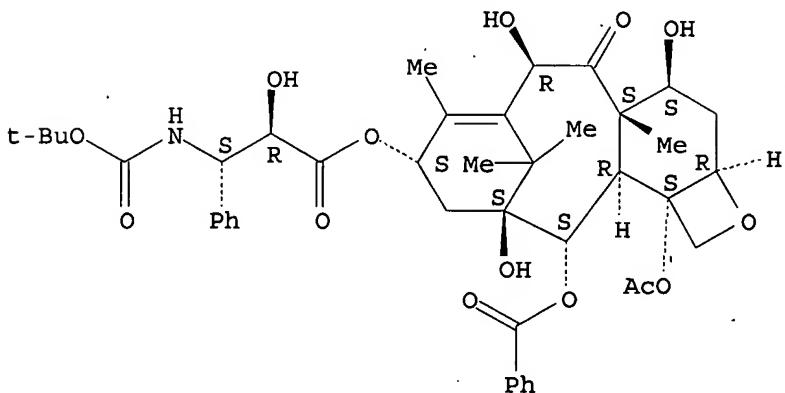
AB Methods are provided to treat or prevent neoplasia disorders in a mammal using a combination of a matrix metalloproteinase inhibitor, an integrin antagonist, and an antineoplastic agent.

IT 114977-28-5, Docetaxel
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(matrix metalloproteinase inhibitor and integrin antagonist in neoplasia treatment)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 12 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:441655 CAPLUS

DOCUMENT NUMBER: 133:68922

TITLE: Method of using a cyclooxygenase-2 inhibitor and a matrix metalloproteinase inhibitor as a combination therapy in the treatment of neoplasia

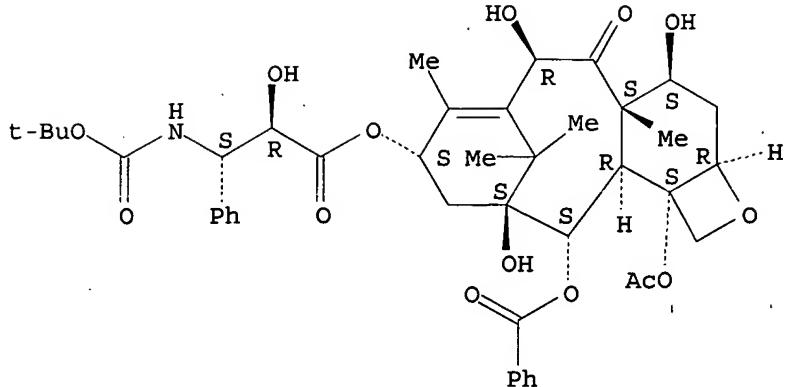
INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.;

Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.
 PATENT ASSIGNEE(S): G.D. Searle and Co., USA
 SOURCE: PCT Int. Appl., 437 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 21
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|-----------------|-------------------|--------------|
| WO 2000037107 | A2 | 20000629 | WO 1999-US30776 | 19991222 <-- |
| WO 2000037107 | A3 | 20010201 | | |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2356426 | AA | 20000629 | CA 1999-2356426 | 19991222 <-- |
| AU 2000025936 | A5 | 20000712 | AU 2000-25936 | 19991222 <-- |
| EP 1140194 | A2 | 20011010 | EP 1999-968540 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| TR 200102499 | T2 | 20011221 | TR 2001-200102499 | 19991222 |
| BR 9916536 | A | 20020102 | BR 1999-16536 | 19991222 |
| HU 200104747 | A2 | 20020429 | HU 2001-4747 | 19991222 |
| JP 2002532563 | T2 | 20021002 | JP 2000-589217 | 19991222 |
| EP 1522313 | A1 | 20050413 | EP 2004-26577 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, RO, CY | | | | |
| AT 322290 | E | 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A | 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A | 20020107 | ZA 2001-5120 | 20010621 |
| NO 2001003156 | A | 20010823 | NO 2001-3156 | 20010622 |
| AU 2004210578 | A1 | 20041007 | AU 2004-210578 | 20040910 |
| PRIORITY APPLN. INFO.: | | | | |
| | | US 1998-113786P | P 19981223 | |
| | | US 1999-385214 | A 19990827 | |
| | | AU 2000-25936 | A3 19991222 | |
| | | EP 1999-968939 | A3 19991222 | |
| | | WO 1999-US30776 | W 19991222 | |

- AB Methods are provided to treat or prevent neoplasia disorders in a mammal using a combination of a cyclooxygenase-2 inhibitor, a matrix metalloproteinase inhibitor and an antineoplastic agent.
 IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (cyclooxygenase-2 inhibitor and matrix metalloproteinase inhibitor in combination therapy for neoplasia treatment)
 RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 13 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1997:400101 CAPLUS

DOCUMENT NUMBER: 127:23742

TITLE: Method, compositions and kits for increasing the oral bioavailability of pharmaceutical agents

INVENTOR(S): Broder, Samuel; Duchin, Kenneth L.; Selim, Sami

PATENT ASSIGNEE(S): Baker Norton Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 136 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 9715269 | A2 | 19970501 | WO 1996-IB1485 | 19961024 <-- |
| WO 9715269 | A3 | 19970731 | | |
| W: AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IL, JP, KE, KG, KP, KR, KZ, LK, LR, LT, LU, LV, MD, MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TT, UA, UZ, VN | | | | |
| RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
| US 5968972 | A | 19991019 | US 1996-608776 | 19960229 <-- |
| US 6245805 | B1 | 20010612 | US 1996-733142 | 19961016 |
| AU 9712056 | A1 | 19970515 | AU 1997-12056 | 19961024 <-- |
| AU 698142 | B2 | 19981022 | | |
| EP 794794 | A1 | 19970917 | EP 1996-943268 | 19961024 <-- |
| EP 794794 | B1 | 20051207 | | |
| R: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE | | | | |
| JP 10509741 | T2 | 19980922 | JP 1997-516449 | 19961024 <-- |
| JP 3361102 | B2 | 20030107 | | |
| BR 9607066 | A | 20021210 | BR 1996-7066 | 19961024 |
| RU 2217135 | C2 | 20031127 | RU 1997-112888 | 19961024 |
| PL 188281 | B1 | 20050131 | PL 1996-321791 | 19961024 |
| AT 311903 | E | 20051215 | AT 1996-943268 | 19961024 |
| ZA 9609001 | A | 19970617 | ZA 1996-9001 | 19961025 <-- |
| NO 9702968 | A | 19970723 | NO 1997-2968 | 19970625 <-- |
| NO 321091 | B1 | 20060313 | | |
| HK 1001960 | A1 | 20060127 | HK 1998-101042 | 19980211 |
| AU 2002035584 | A5 | 20020606 | AU 2002-35584 | 20020422 |
| AU 784159 | B2 | 20060216 | | |
| PRIORITY APPLN. INFO.: | | | US 1995-7071P | P 19951026 |
| | | | US 1996-608776 | A 19960229 |

US 1996-733142 A 19961016
 WO 1996-IB1485 W 19961024
 AU 1998-71300 A3 19980422

AB A method of increasing the bioavailability upon oral administration of a pharmacol. active target agent, particularly an antitumor or antineoplastic agent which exhibits poor or inconsistent oral bioavailability (e.g., paclitaxel, docetaxel or etoposide), comprises the oral co-administration to a mammalian patient of the target agent and an oral bioavailability-enhancing agent (e.g., cyclosporin A, cyclosporin D, cyclosporin F, or ketoconazole). The oral bioavailability-enhancing agents are known to be MDR (P-glycoprotein) inhibitors. The enhancing agent may be administered orally from 0.5-24 h prior to the oral administration of one or more doses of the target agent, substantially simultaneously with the target agent, or both prior to and substantially simultaneously with the target agent. A method of treating mammalian patients suffering from diseases responsive to target agents with poor oral bioavailability, as well as oral dosage forms containing such target agents, combination oral dosage forms containing bioavailability-enhancing agents and target agents kits containing enhancing and target agent dosage forms and dosing information for the co-administration of the same are also disclosed.

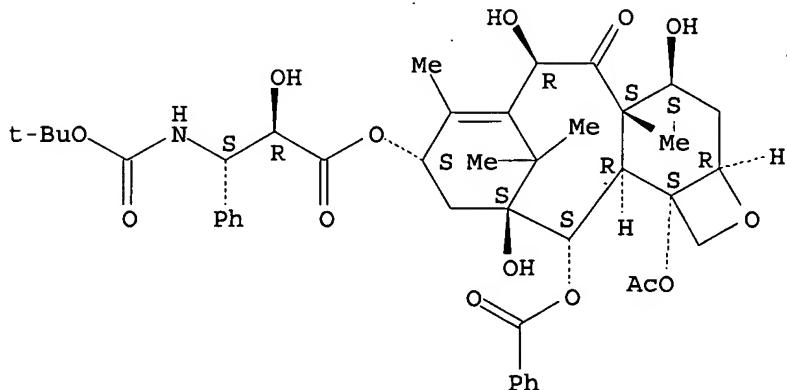
IT 114977-28-5, Docetaxel

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (target; increasing oral bioavailability of pharmaceutical agents)

RN 114977-28-5 CAPLUS

CN Benzene propanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 14 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1998:799989 CAPLUS

DOCUMENT NUMBER: 130:43304

TITLE: Method and compositions for administering taxanes orally to human patients using a cyclosporin to enhance bioavailability

INVENTOR(S): Broder, Samuel; Duchin, Kenneth L.; Selim, Sami

PATENT ASSIGNEE(S): Baker Norton Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 44 pp.

CODEN: PIIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|--------------|
| WO 9853811 | A1 | 19981203 | WO 1998-US7776 | 19980422 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG,
KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
UA, UG, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
| CA 2290446 | AA | 19981203 | CA 1998-2290446 | 19980422 <-- |
| AU 9871300 | A1 | 19981230 | AU 1998-71300 | 19980422 <-- |
| EP 994706 | A1 | 20000426 | EP 1998-918361 | 19980422 <-- |
| EP 994706 | B1 | 20051102 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI, RO | | | | |
| BR 9809694 | A | 20001003 | BR 1998-9694 | 19980422 <-- |
| JP 2002500667 | T2 | 20020108 | JP 1999-500663 | 19980422 |
| HU 200003546 | A2 | 20021128 | HU 2000-3546 | 19980422 |
| RU 2205005 | C2 | 20030527 | RU 1999-128033 | 19980422 |
| NZ 516026 | A | 20030630 | NZ 1998-516026 | 19980422 |
| CN 1550231 | A | 20041201 | CN 2004-10030478 | 19980422 |
| AT 308365 | E | 20051115 | AT 1998-918361 | 19980422 |
| ES 2247690 | T3 | 20060301 | ES 1998-918361 | 19980422 |
| ZA 9804268 | A | 19990623 | ZA 1998-4268 | 19980520 <-- |
| HK 1026637 | A1 | 20060106 | HK 2000-105943 | 20000920 |
| AU 2002035584 | A5 | 20020606 | AU 2002-35584 | 20020422 |
| AU 784159 | B2 | 20060216 | | |

PRIORITY APPLN. INFO.:

| | |
|----------------|-------------|
| US 1997-863513 | A 19970527 |
| AU 1998-71300 | A3 19980422 |
| NZ 1998-501127 | A1 19980422 |
| WO 1998-US7776 | W 19980422 |

AB Taxane antineoplastic agents which have heretofore exhibited poor or non-existent oral bioavailability are administered orally to human patients suffering from taxane-responsive disease conditions and made sufficiently bioavailable to achieve therapeutic blood levels. In a preferred embodiment, the taxane, preferably paclitaxel, is co-administered to the patient with an oral cyclosporin enhancing agent, preferably cyclosporin A. By one preferred method, a dose of oral enhancer is administered about 0.5-72 h before the taxane and a second dose of the enhancer and administered immediately before, together with or immediately after the taxane. A method of treating human patients suffering from taxane-responsive disease conditions is also provided, as well as a method for providing such treatment while preventing or reducing hypersensitivity and allergic reactions without the need for pre-medication.

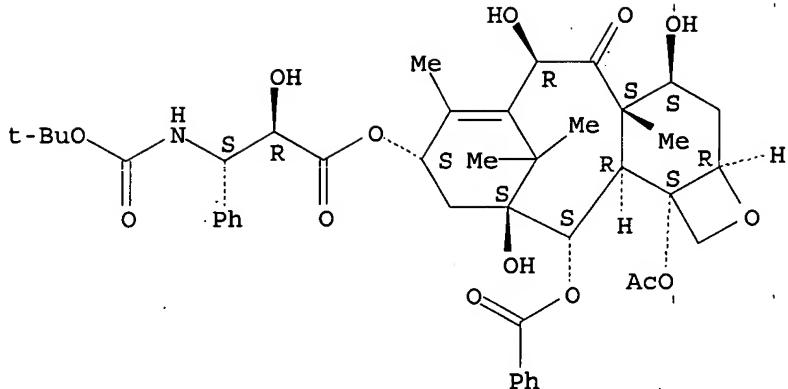
IT 114977-28-5, Docetaxel

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(method and compns. for administering taxanes orally to human patients using a cyclosporin to enhance bioavailability).

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 15 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1999:215571 CAPLUS

DOCUMENT NUMBER: 130:247032

TITLE: Fused imidazole derivatives for improving oral bioavailability of pharmaceutical agents

INVENTOR(S): Snoeck, Henricus Johannes Matheus

PATENT ASSIGNEE(S): Janssen Pharmaceutica N.V., Belg.

SOURCE: PCT Int. Appl., 45 pp.

CODEN: PIXXD2

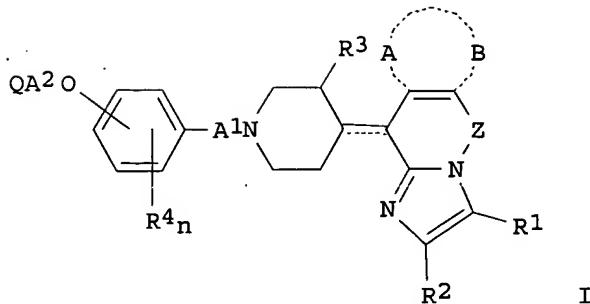
DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|--------|------------|-----------------|--------------|
| WO 9913871 | A2 | 19990325 | WO 1998-EP5751 | 19980910 <-- |
| WO 9913871 | A3 | 19990603 | | |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG,
KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
UA, UG, US, UZ, VN, YU, ZW
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 9911460 | A1 | 19990405 | AU 1999-11460 | 19980910 <-- |
| EP 1011726 | A2 | 20000628 | EP 1998-954268 | 19980910 <-- |
| EP 1011726 | B1 | 20030226 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE,
SI, LT, LV, FI, RO | | | | |
| JP 2001516716 | T2 | 20011002 | JP 2000-511494 | 19980910 |
| JP 3775780 | B2 | 20060517 | | |
| AT 233104 | E | 20030315 | AT 1998-954268 | 19980910 |
| ZA 9808527 | A | 20000322 | ZA 1998-8527 | 19980917 <-- |
| US 6544979 | B1 | 20030408 | US 2000-508748 | 20000315 |
| PRIORITY APPLN. INFO.: | | | EP 1997-202862 | A 19970918 |
| | | | WO 1998-EP5751 | W 19980910 |
| OTHER SOURCE(S): | MARPAT | 130:247032 | | |
| GI | | | | |



AB Compds. I [dotted line = optional bond; n = 1, 2; R1 = H, halo, formyl, (substituted) C1-4 alkyl, etc.; R2 = H, halo, C1-4 alkyl, hydroxy-C1-4 alkyl, etc.; R3 = H, C1-4 alkyl, C1-4 alkyloxy; R4 = H, halo, C1-4 alkyl, C1-4 alkyloxy, halo-C1-4 alkyl; Z = CH₂, CH₂CH₂, CH=CH, CH(OH)CH₂, OCH₂, C(O)CH₂, C(=NOH)CH₂; AB = bivalent radical; A1 = direct bond, (substituted) C1-6 alkanediyl, C1-6 alkanediyl-oxy-C1-6 alkanediyl, carbonyl, C1-6 alkanediylcarbonyl, (substituted) C1-6 alkanediylloxy; A2 = direct bond, C1-6 alkanediyl; Q = aryl], and N-oxide forms, pharmaceutically acceptable addition salts, and stereochem. isomeric forms thereof, are used for the manufacture of a medicine for improving the bioavailability of a second pharmaceutical agent which is co-administered orally to a warm-blooded animal. The second pharmaceutical agent is e.g. an antitumor agent. Preparation of compds. of the invention, and intermediates thereto, is described.

IT 114977-28-5, Docetaxel

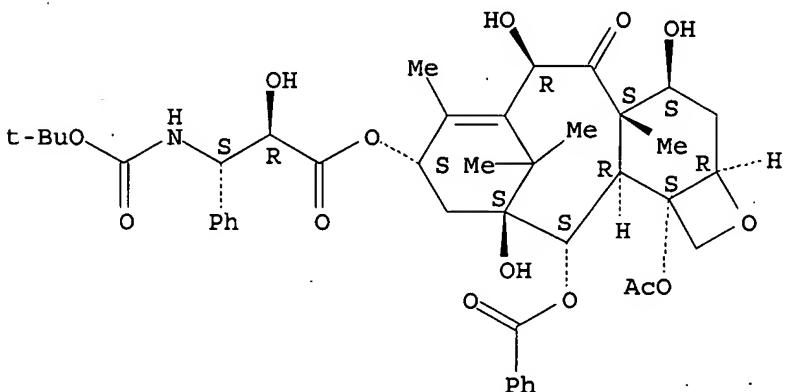
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(fused imidazole derivs., and preparation thereof, for improving oral bioavailability of pharmaceutical agents)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



TITLE: Down-regulation of DNA repair to enhance sensitivity
 to p53-mediated suppression in cancer therapy
 INVENTOR(S): Gjerset, Ruth A.
 PATENT ASSIGNEE(S): Sidney Kimmel Cancer Center, USA; Gjerset, Ruth A.
 SOURCE: PCT Int. Appl., 88 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 9801123 | A1 | 19980115 | WO 1997-US12542 | 19970702 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ,
LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL,
PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US,
UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR,
GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,
GN, ML, MR, NE, SN, TD, TG | | | | |
| US 6054467 | A | 20000425 | US 1996-675887 | 19960705 <-- |
| CA 2259960 | AA | 19980115 | CA 1997-2259960 | 19970702 <-- |
| AU 9736705 | A1 | 19980202 | AU 1997-36705 | 19970702 <-- |
| AU 724212 | B2 | 20000914 | | |
| EP 910357 | A1 | 19990428 | EP 1997-933543 | 19970702 <-- |
| EP 910357 | B1 | 20030604 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI | | | | |
| JP 2000516207 | T2 | 20001205 | JP 1998-505400 | 19970702 <-- |
| AT 241973 | E | 20030615 | AT 1997-933543 | 19970702 |
| US 2005095226 | A1 | 20050505 | US 2004-842718 | 20040510 |
| PRIORITY APPLN. INFO.: | | | US 1996-675887 | A2 19960705 |
| | | | WO 1997-US12542 | W 19970702 |
| | | | US 2000-556440 | B1 20000424 |

AB The present invention details methods for the treatment of cancer. In particular, it concerns the induction of apoptosis in cancer cells following treatment with inhibitors of DNA repair in combination with p53 gene therapy. Treatment of glioblastoma and breast tumor cells with inhibitors of DNA repair induced growth suppression that was a result of p53-mediated apoptosis. Thus it appears that inhibitors of DNA repair in combination with p53 gene therapy is involved in restoration of p53-mediated apoptosis.

IT 114977-28-5, Taxotere

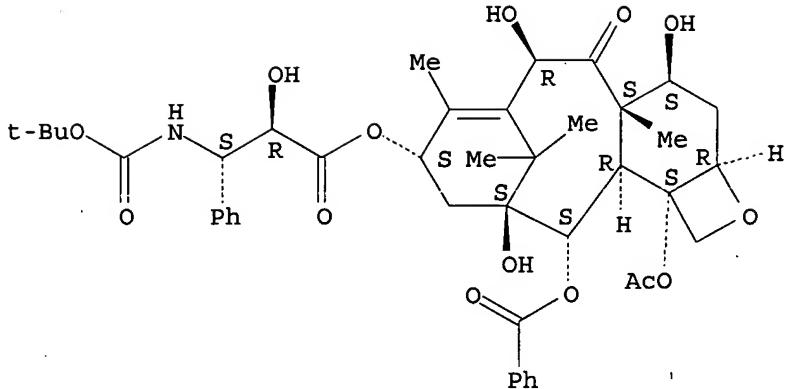
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(down-regulation of DNA repair to enhance sensitivity to p53-mediated suppression in cancer therapy)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S) - (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 17 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1999:405000 CAPLUS

DOCUMENT NUMBER: 131:43591

TITLE: Combination therapy of cancer with anti-ErbB2 antibodies

INVENTOR(S): Shak, Steven; Paton, Virginia E.

PATENT ASSIGNEE(S): Genentech, Inc., USA

SOURCE: PCT Int. Appl., 42 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-------------------|--------------|
| WO 9931140 | A1 | 19990624 | WO 1998-US26266 | 19981210 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE,
KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW,
MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR,
TT, UA, UG, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| ZA 9811162 | A | 20000607 | ZA 1998-11162 | 19981207 <-- |
| CA 2311409 | AA | 19990624 | CA 1998-2311409 | 19981210 <-- |
| AU 9919081 | A1 | 19990705 | AU 1999-19081 | 19981210 <-- |
| EP 1037926 | A1 | 20000927 | EP 1998-963840 | 19981210 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO | | | | |
| TR 200001689 | T2 | 20010122 | TR 2000-200001689 | 19981210 |
| CN 1281468 | A | 20010124 | CN 1998-812097 | 19981210 |
| BR 9815363 | A | 20011016 | BR 1998-15363 | 19981210 |
| JP 2002508397 | T2 | 20020319 | JP 2000-539062 | 19981210 |
| CN 1820734 | A | 20060823 | CN 2006-10008639 | 19981210 |
| NZ 504597 | A | 20030530 | NZ 2000-504597 | 20000517 |
| NO 2000002957 | A | 20000811 | NO 2000-2957 | 20000609 <-- |
| US 2003147884 | A1 | 20030807 | US 2003-356824 | 20030203 |
| US 2004037823 | A9 | 20040226 | | |
| US 2003170234 | A1 | 20030911 | US 2003-406925 | 20030404 |
| US 2005002928 | A1 | 20050106 | US 2004-909998 | 20040802 |
| PRIORITY APPLN. INFO.: | | | US 1997-69346P | P 19971212 |
| | | | CN 1998-812097 | A3 19981210 |
| | | | US 1998-208649 | A3 19981210 |

AB The authors disclose the treatment of disorders characterized by the overexpression of ErbB2. More specifically, human patients are treated with a combination of an anti-ErbB2 antibody and a chemotherapeutic agent other than an anthracycline (e.g., doxorubicin or epirubicin). Preferably, the chemotherapeutic agent is Taxol.

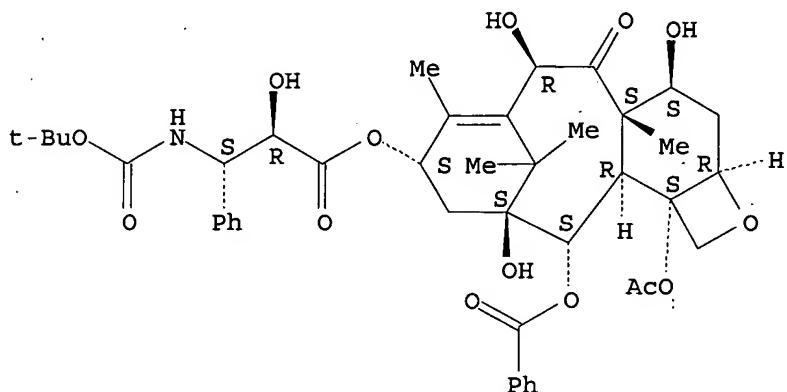
IT 114977-28-5

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (in combination cancer therapy with anti-erbB-2 receptor antibodies)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[(1,1-dimethylethoxy)carbonyl]amino- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 18 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:911036 CAPLUS

DOCUMENT NUMBER: 134:76383

TITLE: Oral pharmaceutical compositions containing taxanes

INVENTOR(S): Gutierrez-Rocca, Jose C.; Cacace, Janice L.; Selim, Sami; Testman, Robert; Rutledge, J. Michael

PATENT ASSIGNEE(S): Baker Norton Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 48 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|-------------|
| WO 2000078247 | A1 | 20001228 | WO 1999-US13821 | 19990618 <- |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |

| | | | |
|--|-------------|-----------------|--------------|
| CA 2371924 | AA 20001228 | CA 1999-2371924 | 19990618 <-- |
| AU 9946955 | A1 20010109 | AU 1999-46955 | 19990618 |
| AU 774060 | B2 20040617 | | |
| BR 9917403 | A 20020709 | BR 1999-17403 | 19990618 |
| EP 1221908 | A1 20020717 | EP 1999-930408 | 19990618 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI, RO, CY | | | |
| JP 2003502349 | T2 20030121 | JP 2001-504316 | 19990618 |
| HU 200300836 | A2 20030828 | HU 2003-836 | 19990618 |
| NZ 516279 | A 20040625 | NZ 1999-516279 | 19990618 |
| RU 2236226 | C2 20040920 | RU 2002-100703 | 19990618 |
| EP 1479382 | A1 20041124 | EP 2004-77062 | 19990618 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI, RO, CY | | | |

PRIORITY APPLN. INFO.: EP 1999-930408 A3 19990618
WO 1999-US13821 W 19990618

AB Pharmaceutical compns. for oral administration to mammalian subjects comprise a taxane or taxane derivative (e.g., paclitaxel or docetaxel) as active ingredient and a vehicle comprising at least 30% by weight of a carrier for the taxane, the carrier having an HLB value of at least about 10. The compns. may also comprise 0-70% of a viscosity-reducing co-solubilizer. The compns. may be incorporated into conventional oral pharmaceutical dosage forms, or can be in the form of a 2-part drug wherein the first part includes the taxane in a solubilizing vehicle and the second part comprises a carrier for the taxane to promote oral absorption. Methods of treatment of taxane-responsive disease conditions employing the novel compns. are also disclosed, whereby the compns. can be administered alone or in association with an oral bioavailability enhancing agent. A formulation containing Tween 80 at 18 mg/kg and paclitaxel gave an absolute bioavailability of 54% which was >15% for i.v. drug.

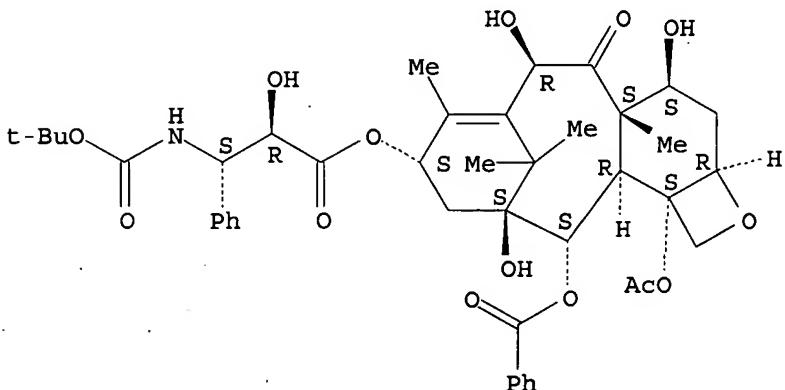
IT 114977-28-5, Docetaxel

RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses) (oral pharmaceuticals containing taxanes)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



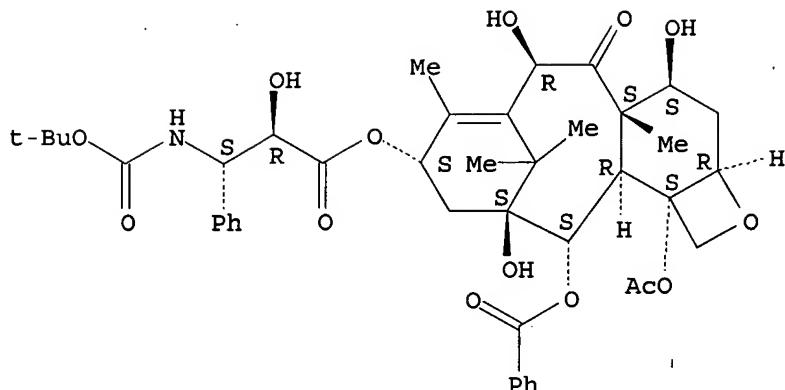
REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ACCESSION NUMBER: 2005:1210172 CAPLUS
 DOCUMENT NUMBER: 143:466194
 TITLE: Oral pharmaceutical compositions containing taxanes
 and methods of cancer therapy employing the same
 INVENTOR(S): Gutierrez-Rocca, Jose C.; Cacace, Janice L.; Selim,
 Sami; Testman, Robert; Rutledge, J. Michael
 PATENT ASSIGNEE(S): Baker Norton Pharmaceuticals, Inc., USA
 SOURCE: U.S., 14 pp., Cont.-in-part of U.S. Ser. No. 863,513,
 abandoned.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 4
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-----------------|-------------|
| US 6964946 | B1 | 20051115 | US 1998-55818 | 19980406 |
| US 5968972 | A | 19991019 | US 1996-608776 | 19960229 <- |
| US 6245805 | B1 | 20010612 | US 1996-733142 | 19961016 |
| ZA 9609001 | A | 19970617 | ZA 1996-9001 | 19961025 <- |
| NZ 516026 | A | 20030630 | NZ 1998-516026 | 19980422 |
| AU 2002035584 | A5 | 20020606 | AU 2002-35584 | 20020422 |
| AU 784159 | B2 | 20060216 | | |
| US 2005267201 | A1 | 20051201 | US 2005-165896 | 20050624 |
| PRIORITY APPLN. INFO.: | | | US 1995-7071P | P 19951026 |
| | | | US 1996-608776 | A2 19960229 |
| | | | US 1996-733142 | A2 19961016 |
| | | | US 1997-863513 | B2 19970527 |
| | | | US 1998-55818 | A3 19980406 |
| | | | AU 1998-71300 | A3 19980422 |
| | | | NZ 1998-501127 | A1 19980422 |

- AB The present invention relates to pharmaceutical compns. for oral administration to mammalian subjects comprising a taxane or taxane derivative (e.g., paclitaxel or docetaxel) as active ingredient and a vehicle comprising at least 30% by weight of a carrier for the taxane, said carrier having an HLB value of at least about 10. The compns. may also comprise 0-70% of a viscosity-reducing co-solubilizer. The compns. may be incorporated into conventional oral pharmaceutical dosage forms, or can be in the form of a two-part medicament wherein the first part includes the taxane in a solubilizing vehicle and the second part comprises a carrier for the taxane to promote oral absorption. Methods of treatment of taxane-responsive disease conditions employing the novel compns. are also disclosed, whereby the compns. can be administered alone or in association with an oral bioavailability enhancing agent.
- IT 114977-28-5, Docetaxel
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (oral pharmaceutical compns. containing taxanes and methods of cancer therapy employing same)
- RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -[(1,1-dimethylethoxy)carbonyl]amino- α -hydroxy-, (2aR,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 20 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:191189 CAPLUS

DOCUMENT NUMBER: 132:227475

TITLE: Treatment of oncologic tumors with an injectable formulation of a Golgi apparatus disturbing agent

INVENTOR(S): Singh, Saira Sayed

PATENT ASSIGNEE(S): Oncopharmaceutical, Inc., USA

SOURCE: PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|-----------------|--------------|
| WO 2000015766 | A1 | 20000323 | WO 1999-US21312 | 19990915 <-- |
| W: AU, CA, JP, KR
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
PT, SE | | | | |
| CA 2344316 | AA | 20000323 | CA 1999-2344316 | 19990915 <-- |
| AU 9959253 | A1 | 20000403 | AU 1999-59253 | 19990915 <-- |
| EP 1114144 | A1 | 20010711 | EP 1999-946955 | 19990915 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI | | | | |
| US 6287602 | B1 | 20010911 | US 1999-397390 | 19990915 |
| JP 2002525268 | T2 | 20020813 | JP 2000-570293 | 19990915 |
| US 2002012703 | A1 | 20020131 | US 2001-912115 | 20010723 |
| US 6497904 | B2 | 20021224 | | |

PRIORITY APPLN. INFO.: US 1998-100479P P 19980916
US 1999-397390 A1 19990915
WO 1999-US21312 W 19990915

AB Novel pharmaceutical formulations for treating a cellular proliferative disease are provided comprising: a therapeutically effective amount of a Golgi apparatus disturbing agent; a biocompatible carrier; and a solvent. In preferred formulations, the Golgi apparatus disturbing agent is brefeldin A (BFA) and the biocompatible carrier is a polymer such as chitin or chitosan. Methods of treating cellular proliferative diseases using the pharmaceutical formulations are also described. Nude mice bearing human epithelial (KB-1) tumors were treated with a BFA/chitin/dimethylacetamide composition

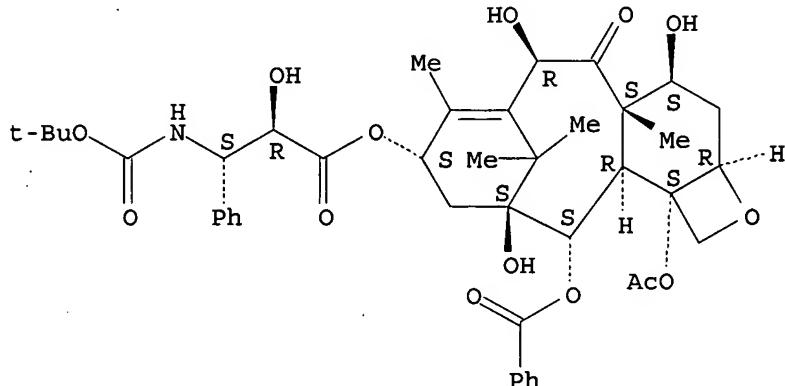
IT 114977-28-5, Docetaxel

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(as addnl. pharmacol. agent; treatment of oncol. tumors with injectable formulation of golgi apparatus disturbing agent)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 21 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:456819 CAPLUS

DOCUMENT NUMBER: 133:84238

TITLE: 3-heteroarylidenyl-2-indolinone compounds for modulating protein kinase activity and for use in cancer chemotherapy

INVENTOR(S): Langecker, Peter J.; Shawver, Laura Kay; Tang, Peng Cho; Sun, Li

PATENT ASSIGNEE(S): Sugen, Inc., USA

SOURCE: PCT Int. Appl., 148 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|-------------|
| WO 2000038519 | A1 | 20000706 | WO 1999-US31232 | 19991230 <- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2357042 | AA | 20000706 | CA 1999-2357042 | 19991230 <- |
| BR 9916735 | A | 20010925 | BR 1999-16735 | 19991230 |
| EP 1139754 | A1 | 20011010 | EP 1999-966725 | 19991230 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| JP 2002533360 | T2 | 20021008 | JP 2000-590484 | 19991230 |
| AU 760964 | B2 | 20030522 | AU 2000-22215 | 19991230 |
| WO 2001049287 | A1 | 20010712 | WO 2000-US18058 | 20000630 |

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 CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
 ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
 LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
 SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
 CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
 EP 1259234 A1 20021127 EP 2000-943334 20000630
 EP 1259234 B1 20060816
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL
 JP 2003535038 T2 20031125 JP 2001-549655 20000630
 AT 336245 E 20060915 AT 2000-943334 20000630
 US 2003191162 A1 20031009 US 2002-307483 20021202
 PRIORITY APPLN. INFO.: US 1998-114313P P 19981231
 US 1999-476232 A 19991230
 WO 1999-US31232 W 19991230
 US 2000-569545 A 20000512
 WO 2000-US18058 W 20000630

OTHER SOURCE(S): MARPAT 133:84238

AB 3-Heteroarylidenyl-2-indolinone compds. are provided that modulate the enzymic activity of protein kinases and therefore are expected to be useful in the prevention and treatment of protein kinase-related cellular disorders, e.g. cancer. Furthermore, these compds. are expected to enhance the efficacy of other chemotherapeutic agents, in particular, fluorinated pyrimidines, in the treatment of cancer.

IT 114977-28-5, Docetaxel

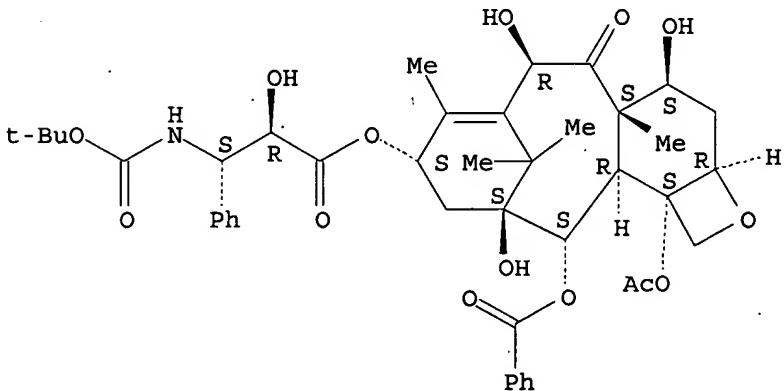
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(heteroarylidenylindolinone derivs. for modulating protein kinase activity and in cancer chemotherapy)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 22 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2000:880923 CAPLUS

DOCUMENT NUMBER: 134:37055
 TITLE: Methods and compositions using FGF inhibitors and
 agonists for modulating cell proliferation and cell
 death
 INVENTOR(S): Au, Jessie L. S.; Wientjes, M. Guillaume
 PATENT ASSIGNEE(S): USA
 SOURCE: PCT Int. Appl., 143 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|-----------------|-----------------|--------------|
| WO 2000074634 | A2 | 20001214 | WO 2000-US40103 | 20000605 <-- |
| WO 2000074634 | C2 | 20020926 | | |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID,
IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV,
MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE,
SG, SI, SK | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2377385 | AA | 20001214 | CA 2000-2377385 | 20000605 <-- |
| EP 1206234 | A2 | 20020522 | EP 2000-943429 | 20000605 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL | | | | |
| JP 2003503313 | T2 | 20030128 | JP 2001-501171 | 20000605 |
| US 6599912 | B1 | 20030729 | US 2000-587559 | 20000605 |
| AU 780454 | B2 | 20050324 | AU 2000-57903 | 20000605 |
| US 2004010001 | A1 | 20040115 | US 2003-464018 | 20030618 |
| PRIORITY APPLN. INFO.: | | | | |
| | | US 1999-137345P | P 19990603 | |
| | | US 1999-165983P | P 19991117 | |
| | | US 1999-172031P | P 19991223 | |
| | | US 2000-187445P | P 20000307 | |
| | | US 2000-587559 | A3 20000605 | |
| | | WO 2000-US40103 | W 20000605 | |

AB Methods and compns. for modulating the FGF effect on the sensitivity of malignant and normal cells to anticancer agents are provided. In particular, methods and compns. for inhibiting FGF-induced resistance to a broad spectrum of anticancer agents in solid and soft-tissue tumors, metastatic lesions, leukemia and lymphoma are provided. Preferably, the compns. include at least one FGF inhibitor in combination with a cytotoxic agents, e.g., antimicrotubule agents, topoisomerase I inhibitors, topoisomerase II inhibitors, antimetabolites, mitotic inhibitors, alkylating agents, intercalating agents, agents capable of interfering with a signal transduction pathway (e.g., g., a protein kinase C inhibitor, e.g., an anti-hormone, e.g., an antibody against growth factor receptors), an agent that promote apoptosis and/or necrosis, an interferon, an interleukin, a tumor necrosis factor, and radiation. In other embodiments, methods and composition for protecting a cell in a subject, from one or more of killing, inhibition of growth or division or other damage caused, e.g., by a cytotoxic agent, are provided. Preferably, the method includes administering to the subject an effective amount of at least one FGF agonist, thereby treating the cell, e.g., protecting or reducing the damage to the dividing cell from said cytotoxic agent. FGF gene expression-based methods for diagnosis of proliferative disorders are also disclosed.

IT 114977-28-5, Taxotere

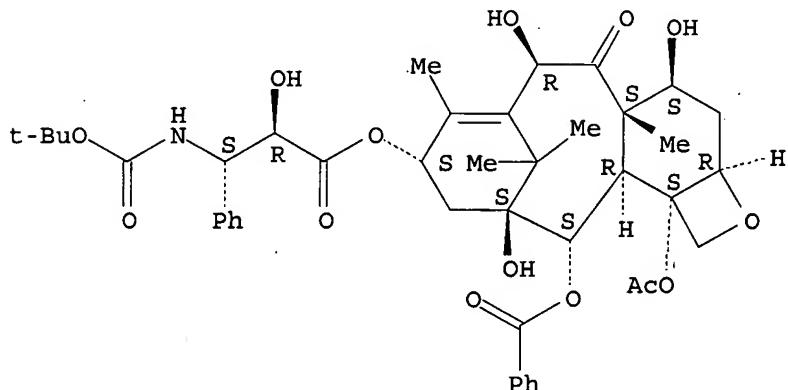
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(FGF inhibitors and agonists for modulating cell proliferation and cell death)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy) carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 23 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN.

ACCESSION NUMBER: 2000:824125 CAPLUS

DOCUMENT NUMBER: 134:4050

TITLE: Treatment with anti-erbB2 antibodies

INVENTOR(S): Cohen, Robert L.

PATENT ASSIGNEE(S): Genentech, Inc., USA

SOURCE: PCT Int. Appl., 39 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 2000069460 | A1 | 20001123 | WO 2000-US12552 | 20000509 <-- |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2374085 | AA | 20001123 | CA 2000-2374085 | 20000509 <-- |
| EP 1187632 | A1 | 20020320 | EP 2000-928916 | 20000509 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| JP 2002544238 | T2 | 20021224 | JP 2000-617920 | 20000509 |
| AU 782325 | B2 | 20050721 | AU 2000-47080 | 20000509 |
| US 2003170235 | A1 | 20030911 | US 2003-429519 | 20030505 |
| PRIORITY APPLN. INFO.: | | | US 1999-134085P | P 19990514 |
| | | | US 2000-568322 | A1 20000509 |
| | | | WO 2000-US12552 | W 20000509 |

AB A method treating a human patient to or diagnosed with a tumor in which erbB2 protein is expressed comprising the following steps, performed sequentially: (a) treating the patient with a therapeutically effective amount of an anti-erbB2 antibody; (b) surgically removing the tumor, and then (c) treating the patient with a therapeutically effective amount of an anti-erbB2 antibody or of a chemotherapeutic agent.

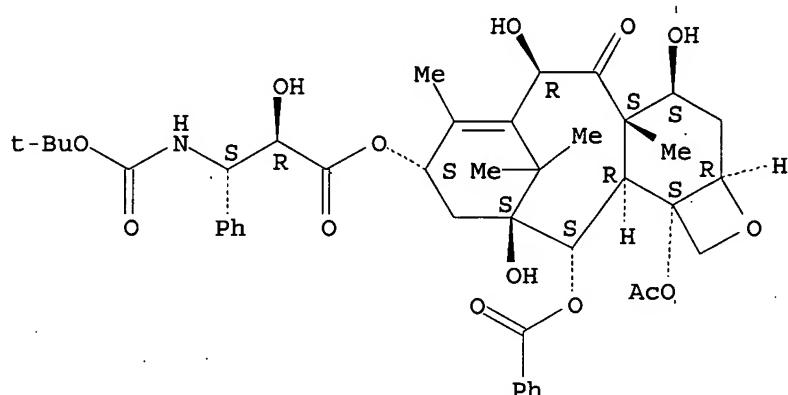
IT 114977-28-5

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(cancer treatment with anti-erbB2 antibodies)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy) carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 24 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:158387 CAPLUS

DOCUMENT NUMBER: 136:210551

TITLE: Method of treating hyperproliferative diseases using active vitamin D analogues

INVENTOR(S): Bishop, Charles W.; Mazess, Richard B.

PATENT ASSIGNEE(S): Bone Care International, Inc., USA

SOURCE: U.S. Pat. Appl. Publ., 13 pp., Cont.-in-part of U.S. Ser. No. 596,149.

CODEN: USXXCO

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC.. NUM. COUNT: 20

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---------------|------|----------|-----------------|-------------|
| US 2002025950 | A1 | 20020228 | US 2001-891814 | 20010626 |
| US 6503893 | B2 | 20030107 | | |
| US 5763429 | A | 19980609 | US 1996-781910 | 19961230 <- |
| US 6537982 | B1 | 20030325 | US 1998-596149 | 19980223 |
| US 2002128240 | A1 | 20020912 | US 2001-995911 | 20011128 |
| CA 2450942 | AA | 20030103 | CA 2002-2450942 | 20020626 |
| WO 2003000023 | A2 | 20030103 | WO 2002-US20475 | 20020626 |
| WO 2003000023 | A3 | 20030731 | | |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,

CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
 GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
 LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
 PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
 UA, UG, UZ, VN, YU, ZA, ZM, ZW
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
 KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB,
 GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA,
 GN, GQ, GW, ML, MR, NE, SN, TD, TG
 EP 1408983 A2 20040421 EP 2002-756332 20020626
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, FI, CY, TR
 CN 1520302 A 20040811 CN 2002-812881 20020626
 JP-2004535429 T2 20041125 JP 2003-506479 20020626
 US 2003130242 A1 20030710 US 2003-337506 20030107
 US 6680309 B2 20040120
 PRIORITY APPLN. INFO.:
 US 1996-781910 A3 19961230
 US 1998-596149 A2 19980223
 US 1993-119895 A2 19930910
 US 1994-265438 A2 19940624
 US 1995-415488 A2 19950403
 US 1995-486387 A2 19950607
 US 2001-891814 A2 20010626
 WO 2002-US20475 W 20020626

OTHER SOURCE(S): MARPAT 136:210551

AB Methods use hypocalcemic vitamin D analogs to inhibit the hyperproliferation of malignant or neoplastic cells without incidence of hypercalcemia. Patients with advanced androgen-independent prostate cancer were treated with 1 α ,24-dihydroxyvitamin D2.

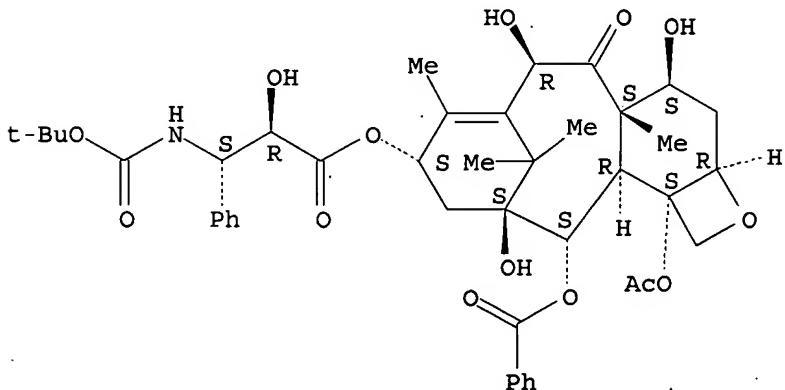
IT 114977-28-5, Docetaxel

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (coadministration with cytotoxic; hypocalcemic vitamin D analogs for treating hyperproliferative diseases)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy) carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



TITLE: Treatment of patients with liver metastases
AUTHOR(S): Fumoleau, P.
CORPORATE SOURCE: Center Regionale de Lutte Contre le Cancer,
Nantes-Atlantique, Herblain, 44805, Fr.
SOURCE: Anti-Cancer Drugs (1996), 7(Suppl. 2,
Management of Advanced Breast Cancer: Patient Needs,
Challenges and New Treatment Options), 21-23
CODEN: ANTDEV; ISSN: 0959-4973

PUBLISHER: Rapid Science Publishers
DOCUMENT TYPE: Journal
LANGUAGE: English

AB The presence of liver metastases is a very poor prognostic factor for patients with metastatic breast cancer. Liver metastases are generally less responsive to chemotherapy than metastases in other sites, and patients with liver lesions have a shorter survival duration than patients with other sites of disease. The results from 5 multicenter phase II studies of docetaxel as a first-line treatment for metastatic breast cancer were analyzed with regard to the presence or absence of liver lesions, which were found in 39% of the 209 patients involved. Response rates to docetaxel, 100 or 75 mg/m², were maintained in the presence of liver lesions and the median survival across all five studies was 16.4 mo for all patients and 14.7 mo for patients with liver lesions. Similarly, when results from 129 patients given docetaxel as a second-line treatment were analyzed, the response rates and survival durations were not reduced in the 57% of patients who had liver lesions. The presence of liver metastases does not reduce the probability or duration of response to docetaxel as a first- or second-line treatment for advanced breast cancer.

IT 114977-28-5, Docetaxel

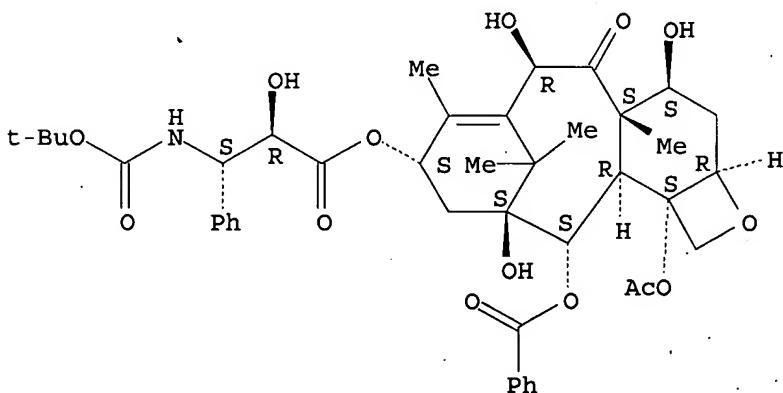
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(treatment of patients with liver metastases)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 26 OF 55 MEDLINE on STN

ACCESSION NUMBER: 2000024224 MEDLINE

DOCUMENT NUMBER: PubMed ID: 10560434

TITLE: A case of multiple liver metastases from breast cancer

successfully treated with intra-arterial administration of docetaxel.
 AUTHOR: Maeda Y; Nishida M; Takao T; Harada K; Mori N; Tamesa T;
 Somura H; Tangoku A; Oka M; Konishi T
 CORPORATE SOURCE: Dept. of Surgery II, Yamaguchi University School of Medicine.
 SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1999 Oct) Vol. 26, No. 12, pp. 1951-4.
 Journal code: 7810034. ISSN: 0385-0684.
 PUB. COUNTRY: Japan
 DOCUMENT TYPE: (CASE REPORTS)
 LANGUAGE: Japanese
 FILE SEGMENT: Priority Journals
 ENTRY MONTH: 199911
 ENTRY DATE: Entered STN: 13 Jan 2000
 Last Updated on STN: 13 Jan 2000
 Entered Medline: 26 Nov 1999

AB Docetaxel is an excellent agent with a high antitumor effect for the treatment of advanced/recurrent breast cancer. A 55-year-old female with metastatic liver tumors from breast cancer showed a remarkable response to intra-arterial administration of docetaxel (20 mg/week, or 40 mg/2 weeks). Since CT and MRI imaging revealed multiple metastases in the liver, intra-arterial chemotherapy was selected. No critical side effect was found during this chemotherapy. A CT scan 3 months after chemotherapy showed a partial response. We conclude that this intra-arterial chemotherapy using docetaxel will be safe and useful for liver metastases from breast cancer.

L12 ANSWER 27 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1998:564267 CAPLUS
 DOCUMENT NUMBER: 129:197984
 TITLE: Combined tumor suppressor gene therapy and chemotherapy in the treatment of neoplasms
 INVENTOR(S): Nielsen, Loretta; Horowitz, Jo Ann; Maneval, Daniel C.; Demers, G. William; Rybak, Mary Ellen; Resnick, Gene
 PATENT ASSIGNEE(S): Canji, Inc., USA
 SOURCE: PCT Int. Appl., 114 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 9835554 | A2 | 19980820 | WO 1998-US3514 | 19980217 <-- |
| WO 9835554 | A3 | 19981126 | | |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
| CA 2282683 | AA | 19980820 | CA 1998-2282683 | 19980217 <-- |
| AU 9864380 | A1 | 19980908 | AU 1998-64380 | 19980217 <-- |
| AU 737621 | B2 | 20010823 | | |
| EP 969720 | A2 | 20000112 | EP 1998-910038 | 19980217 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI | | | | |

| | | | | |
|---------------|----|----------|----------------|--------------|
| NZ 337283 | A | 20010223 | NZ 1998-337283 | 19980217 |
| HU 200004326 | A2 | 20010228 | HU 2000-4326 | 19980217 |
| JP 2001511815 | T2 | 20010814 | JP 1998-536033 | 19980217 |
| BR 9807418 | A | 20020122 | BR 1998-7418 | 19980217 |
| US 2003060434 | A1 | 20030327 | US 1999-311772 | 19990513 |
| NO 9903943 | A | 19991015 | NO 1999-3943 | 19990817 <-- |
| US 2003064949 | A1 | 20030403 | US 2002-86294 | 20020228 |
| US 2004235736 | A1 | 20041125 | US 2004-824058 | 20040413 |
| US 2005142112 | A1 | 20050630 | US 2004-823932 | 20040413 |

PRIORITY APPLN. INFO.:

| | |
|----------------|-------------|
| US 1997-38065P | P 19970218 |
| US 1997-801285 | A 19970218 |
| US 1997-801681 | A 19970218 |
| US 1997-801755 | A 19970218 |
| US 1997-801765 | A 19970218 |
| US 1997-47834P | P 19970528 |
| US 1998-24932 | B1 19980217 |
| WO 1998-US3514 | W 19980217 |
| US 1999-311772 | B3 19990513 |

AB In one embodiment, the invention provides methods of treating mammalian cancer or hyperproliferative cells, the method comprising contacting the cells with a tumor suppressor protein or tumor suppressor nucleic acid and also contacting the cells with at least one adjunctive anticancer agent. The invention also provides for a pharmacol. composition comprising a tumor suppressor protein or a tumor suppressor nucleic acid and at least one adjunctive anti-cancer agent, as well as a kit for the treatment of mammalian cancer or hyperproliferative cells.

IT 114977-28-5, Taxotere

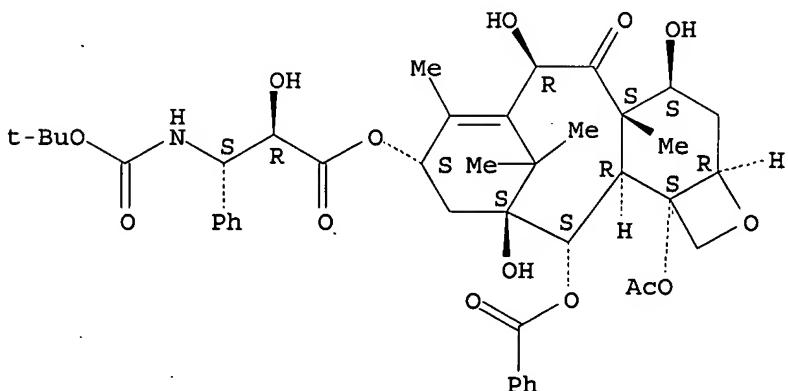
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tumor suppressor gene therapy-chemotherapy combination for treatment of neoplasms and hyperproliferative cells)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 28 OF 55 MEDLINE on STN

ACCESSION NUMBER: 2001066728 MEDLINE

DOCUMENT NUMBER: PubMed ID: 10907946

TITLE: Phase II study of docetaxel in patients with liver metastases from breast cancer. UK study group.

AUTHOR: Coleman R E; Howell A; Eggleton S P; Maling S J; Miles D W
CORPORATE SOURCE: Weston Park Hospital NHS Trust, Sheffield, UK.
SOURCE: Annals of oncology : official journal of the European Society for Medical Oncology / ESMO, (2000 May)
Vol. 11, No. 5, pp. 541-6.
Journal code: 9007735, ISSN: 0923-7534.

PUB. COUNTRY: Netherlands
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
(MULTICENTER STUDY)

LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200012
ENTRY DATE: Entered STN: 22 Mar 2001
Last Updated on STN: 22 Mar 2001
Entered Medline: 28 Dec 2000

AB BACKGROUND: Previous phase II studies of docetaxel have indicated that hepatic metastases from breast cancer respond well to first-line treatment with docetaxel. The objective of this prospective, open label phase II study therefore was specifically to evaluate the activity and safety of docetaxel in this indication. PATIENTS AND METHODS: The study recruited 47 women (mean age 50 years, range 33-66 years) with hepatic metastases from breast cancer who fulfilled the eligibility criteria. After premedication with steroids, patients received a one-hour intravenous infusion of docetaxel 100 mg/m² at three-weekly intervals for up to eight cycles. Response to treatment during medication was assessed after three, six and where appropriate, eight cycles and every three month follow-up thereafter, until disease progression or death. RESULTS: The best overall response rate (ORR) for evaluable patients was 64.3% (95% CI: 48.0-78.5%). In terms of the primary efficacy parameters, the ORR at the sixth cycle of treatment was 62% (95% CI: 45%-80%) with 17% complete responses. The median duration of response was 139 days (95% CI: 111-216 days) and the median survival duration calculated on an intent-to-treat basis was 335 days (227-568 days, 95% CI). One (2%) toxic death was reported. CONCLUSIONS: Docetaxel is a highly effective cytotoxic agent in the treatment of patients with liver metastases from breast cancer.

L12 ANSWER 29 OF 55 MEDLINE on STN
ACCESSION NUMBER: 1998056505 MEDLINE
DOCUMENT NUMBER: PubMed ID: 9446016
TITLE: [Docetaxel (taxotere) for therapy of breast carcinoma. Highest effectiveness with moderate side effects].
Docetaxel (Taxotere) zur Therapie des Mammakarzinoms. Hochste Wirksamkeit bei moderaten Nebenwirkungen.
AUTHOR: von Minckwitz G; Costa S D
CORPORATE SOURCE: Klinik fur Gynakologie und Geburtshilfe, Johann Wolfgang Goethe-Universitat Frankfurt.. minckwitz@em.uni-frankfurt.de
SOURCE: Medizinische Klinik (Munich, Germany : 1983), (1997 Sep 15) Vol. 92 Suppl 4, pp. 4-9. Ref: 16
Journal code: 8303501. ISSN: 0723-5003.
PUB. COUNTRY: GERMANY: Germany, Federal Republic of
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)
General Review; (REVIEW)
LANGUAGE: German
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199801
ENTRY DATE: Entered STN: 6 Feb 1998
Last Updated on STN: 6 Feb 1998
Entered Medline: 27 Jan 1998

AB CLINICAL RESULTS: Docetaxel is a taxan which has proven high efficacy in the treatment of breast cancer. The results are consistent throughout all phases of clinical evaluation. High response rates have been observed especially for women after failure of anthracyclins or with liver metastases. Response rates are superior to doxorubicin, while the extent of the side effects is comparable. CONCLUSION: Due to the different toxicity profile a combination of docetaxel and anthracyclins is feasible and has already been demonstrated in early clinical trials. The role of the combinatory treatments in first line or adjuvant setting is currently under investigation.

L12 ANSWER 30 OF 55 MEDLINE on STN
ACCESSION NUMBER: 96351131 MEDLINE
DOCUMENT NUMBER: PubMed ID: 8745348
TITLE: Docetaxel: a new defence in the management of breast cancer.
AUTHOR: Piccart M
CORPORATE SOURCE: Department of Chemotherapy, Institut Jules Bordet, Brussels, Belgium.
SOURCE: Anti-cancer drugs, (1995 Jul) Vol. 6 Suppl 4, pp. 7-11. Ref: 12
PUB. COUNTRY: Journal code: 9100823. ISSN: 0959-4973.
DOCUMENT TYPE: ENGLAND: United Kingdom
Journal; Article; (JOURNAL ARTICLE)
General Review; (REVIEW)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199610
ENTRY DATE: Entered STN: 25 Oct 1996
Last Updated on STN: 25 Oct 1996
Entered Medline: 16 Oct 1996

AB The results of nine phase II trials of docetaxel in the first- and second-line treatment of patients with advanced breast cancer are summarized. All 316 patients included in this report received docetaxel at a dose of 100 mg/m² administered over 1 h every 3 weeks on an outpatient basis. One hundred and fifty-four patients received docetaxel as first-line therapy for advanced disease, half of whom had received prior adjuvant chemotherapy (finished at least 1 year previously). An overall response rate of 59% (95% CI: 51-67) was achieved in these patients, with a median duration of response of 8.3 months and a median time to progression of 4.9 months. Similar results were seen in a subgroup of 68 patients with liver metastases. Among the 162 patients given docetaxel as second-line therapy, 134 had strictly defined anthracycline-resistant disease; 73 had liver metastases. The combined overall response rate for anthracycline-resistant patients in two US studies was 48% (95% CI: 37-59) while that in a multicenter French study was 29% (95% CI: 18-44). The median duration of response in each case was 6.3 and 5.5 months, respectively, with an overall median survival duration of 11 and 10 months, respectively. Among patients with liver metastases, second-line treatment with docetaxel achieved an overall response rate of 32%, a median duration of response of 7.8 months and a median survival duration of 9 months. These results for docetaxel as both first- and second-line therapy are comparable with those achieved with doxorubicin and are particularly promising in patients with liver metastases and anthracycline-resistant disease.

L12 ANSWER 31 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 1999:736476 CAPLUS
DOCUMENT NUMBER: 131:346535
TITLE: Use of neomycin for treating angiogenesis-related diseases
INVENTOR(S): Hu, Guo-Fu; Vallee, Bert L.
PATENT ASSIGNEE(S): The Endowment for Research In Human Biology, Inc., USA
SOURCE: PCT Int. Appl., 74 pp.

CODEN: PIXXD2

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|-----------------|--------------|
| WO 9958126 | A1 | 19991118 | WO 1999-US10269 | 19990511 <-- |
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JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,
MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2331620 | AA | 19991118 | CA 1999-2331620 | 19990511 <-- |
| AU 9939804 | A1 | 19991129 | AU 1999-39804 | 19990511 <-- |
| EP 1083896 | A1 | 20010321 | EP 1999-922915 | 19990511 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI | | | | |
| US 6482802 | B1 | 20021119 | US 2000-700436 | 20001109 |
| PRIORITY APPLN. INFO.: | | | US 1998-84921P | P 19980511 |
| | | | WO 1999-US10269 | W 19990511 |

AB The present invention is directed to using neomycin or an analog thereof as a therapeutic agent to treat angiogenesis-related diseases, which are characterized by excessive, undesired or inappropriate angiogenesis or proliferation of endothelial cells. The present invention is also directed to pharmaceutical compns. comprising: (a) neomycin or an analog and, optionally, (b) another anti-angiogenic agent or an anti-neoplastic agent.. The present invention is further directed to a method for screening neomycin analogs having anti-angiogenic activity. A preferred embodiment of the invention relates to using neomycin to treat subjects having such diseases. A dose of 20 ng neomycin/embryo or higher completely inhibited angiogenin-induced angiogenesis in the chorioallantoic membrane (CAM) assay. Neomycin inhibits angiogenin-induced angiogenesis mainly through inhibition of nuclear translocation of angiogenin.

IT 114977-28-5, Docetaxel

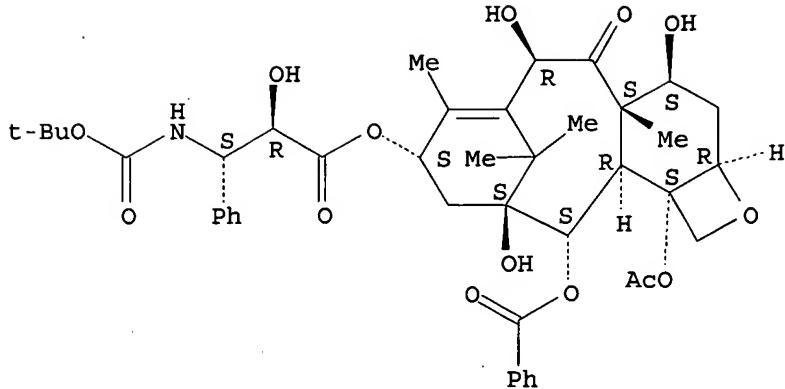
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(neomycin, its analogs and other agents for treatment of angiogenesis-related diseases)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 32 OF 55 MEDLINE on STN

ACCESSION NUMBER: 1999314769 MEDLINE

DOCUMENT NUMBER: PubMed ID: 10408850

TITLE: Phase II study of docetaxel in patients with metastatic pancreatic cancer: a Japanese cooperative study. Cooperative Group of Docetaxel for Pancreatic Cancer in Japan.

AUTHOR: Okada S; Sakata Y; Matsuno S; Kurihara M; Sasaki Y; Ohashi Y; Taguchi T

CORPORATE SOURCE: Department of Internal Medicine, National Cancer Center Hospital, Tokyo, Japan.

SOURCE: British journal of cancer, (1999 May) Vol. 80, No. 3-4, pp. 438-43.

Journal code: 0370635. ISSN: 0007-0920.

PUB. COUNTRY: SCOTLAND: United Kingdom

DOCUMENT TYPE: (CLINICAL TRIAL)

(CLINICAL TRIAL, PHASE II)

Journal; Article; (JOURNAL ARTICLE)

(MULTICENTER STUDY)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199907

ENTRY DATE: Entered STN: 27 Jul 1999

Last Updated on STN: 27 Jul 1999

Entered Medline: 15 Jul 1999

AB Docetaxel has been reported to show promising anti-tumour activity in pancreatic ductal cancer (PC). This study was conducted to evaluate the activity and toxicity of moderate-dose (60 mg m⁻²) docetaxel in Japanese chemo-naive patients with measurable metastatic PC. The patients had a performance status of 0-2. They received docetaxel intravenously over a 1- to 2-h period without any premedication for hypersensitivity reactions. This treatment was repeated every 3-4 weeks with dose adjustments based on the toxic effects observed. Twenty-one patients were eligible and treated with docetaxel. The median number of courses was 2 (range, 1-4). None of the patients achieved an objective response; seven showed no change and 13 showed progressive disease. In one patient, the response was not assessable because of early death. The median survival time for all patients was 118 days. The main grade 3-4 toxicities by patient were leucocytopenia (67%) and neutropenia (86%). Other grade 3-4 toxicities included anaemia (10%), thrombocytopenia (5%), nausea/vomiting (29%), anorexia (29%), GOT/GPT increase (10%), alkaline phosphatase increase (14%), malaise/fatigue (33%) and alopecia (24%). In conclusion, docetaxel, administered on this schedule, did not show significant anti-tumour activity in patients with metastatic PC.

L12 ANSWER 33 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2000:824124 CAPLUS
 DOCUMENT NUMBER: 134:506
 TITLE: Treatment of refractory human tumors with epidermal growth factor receptor antagonists
 INVENTOR(S): Waksal, Harlan W.
 PATENT ASSIGNEE(S): Imclone Systems Incorporated, USA
 SOURCE: PCT Int. Appl., 31 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|--------------|
| WO 2000069459 | A1 | 20001123 | WO 2000-US11756 | 20000501 <-- |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2373815 | AA | 20001123 | CA 2000-2373815 | 20000501 <-- |
| BR 2000010524 | A | 20020528 | BR 2000-10524 | 20000501 |
| EP 1218032 | A1 | 20020703 | EP 2000-928671 | 20000501 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL | | | | |
| HU 200201480 | A2 | 20020828 | HU 2002-1480 | 20000501 |
| EE 200100603 | A | 20030217 | EE 2001-603 | 20000501 |
| JP 2003520195 | T2 | 20030702 | JP 2000-617919 | 20000501 |
| AU 782994 | B2 | 20050915 | AU 2000-46871 | 20000501 |
| CN 1720994 | A | 20060118 | CN 2005-10055865 | 20000501 |
| US 2002012663 | A1 | 20020131 | US 2001-840146 | 20010424 |
| NO 2001005546 | A | 20020114 | NO 2001-5546 | 20011113 |
| ZA 2001009347 | A | 20030213 | ZA 2001-9347 | 20011113 |
| BG 106110 | A | 20020430 | BG 2001-106110 | 20011114 |
| US 2003157104 | A1 | 20030821 | US 2001-996954 | 20011130 |
| US 2005112120 | A1 | 20050526 | US 2004-18950 | 20041220 |
| PRIORITY APPLN. INFO.: | | | US 1999-312284 | A 19990514 |
| | | | US 1999-374028 | A 19990813 |
| | | | CN 2000-810321 | A3 20000501 |
| | | | WO 2000-US11756 | W 20000501 |
| | | | US 2001-840146 | A1 20010424 |

AB A method of inhibiting the growth of refractory tumors that are stimulated by a ligand of epidermal growth factor in human patients comprises treating the human patients with an effective amount of an epidermal growth factor receptor antagonist, e.g. a monoclonal antibody.

IT 114977-28-5, Docetaxel

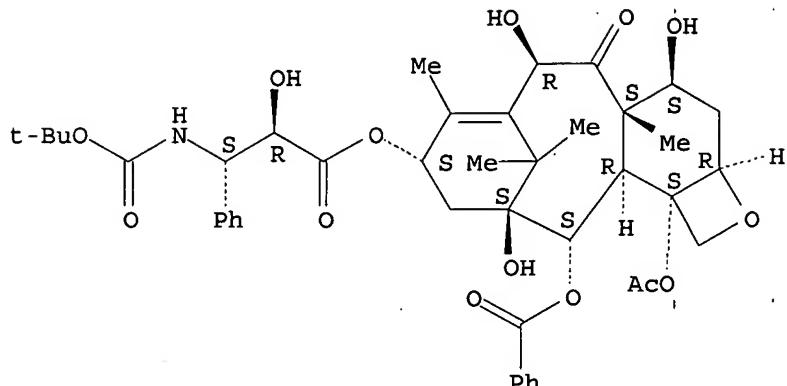
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(EGF receptor antagonists for treatment of refractory human tumors)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 34 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1999:454255 CAPLUS

DOCUMENT NUMBER: 131:92524

TITLE: Therapeutic liposome-encapsulated immunomodulators

INVENTOR(S): Spitler, Lynn E.; Fidler, Issaiah J.

PATENT ASSIGNEE(S): Jenner Biotherapies, Inc., USA

SOURCE: PCT Int. Appl., 111 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 9935162 | A1 | 19990715 | WO 1999-US272 | 19990106 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP,
KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,
MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 9922141 | A1 | 19990726 | AU 1999-22141 | 19990106 <-- |
| US 2003017976 | A1 | 20030123 | US 2001-764546 | 20010117 |
| US 2004146552 | A1 | 20040729 | US 2003-705618 | 20031110 |
| PRIORITY APPLN. INFO.: | | | US 1998-70717P | P 19980107 |
| | | | US 1999-226075 | B1 19990106 |
| | | | WO 1999-US272 | W 19990106 |
| | | | US 2001-764546 | A1 20010117 |

AB The present invention relates to the use of novel compns. of lipopeptides that are immunomodulators encapsulated as liposomes or free-form for the treatment of neoplasia and in reducing chemotherapeutically induced cellular pathol., including mucositis. These lipopeptides may be administered alone or in combination with a second antineoplastic agent. E.g., a synthetic JBT 3002 lipopeptide entrapped in phosphatidylcholine/phosphatidylserine liposomes is shown to be a potent activator of tumoricidal properties of murine macrophages by a mechanism that differs from that of lipopolysaccharides. These data highly support the in vivo use of multilamellar liposome-encapsulated JBT 3002 to enhance host resistance to infections and cancer.

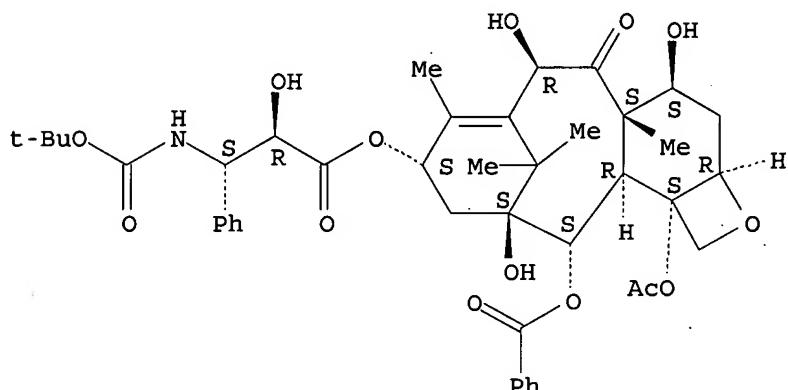
IT 114977-28-5, Taxotere

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combination with; free or liposome-encapsulated lipopeptide immunomodulators for tumor treatment and reduction of antitumor adverse effects)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 35 OF 55 MEDLINE on STN

ACCESSION NUMBER: 97356889 MEDLINE

DOCUMENT NUMBER: PubMed ID: 9213325

TITLE: Docetaxel combined with vinorelbine: phase I results and new study designs.

AUTHOR: Fumoleau P; Fety R; Delecroix V; Perrocheau G; Azli N

CORPORATE SOURCE: Medical Oncology Department, Centre Rene Gauducheau, CRLCC Nantes-Atlantique, Nantes-St Herblain, France.

SOURCE: Oncology (Williston Park, N.Y.), (1997 Jun) Vol. 11, No. 6 Suppl 6, pp. 29-31.

Journal code: 8712059. ISSN: 0890-9091.

PUB. COUNTRY: United States

DOCUMENT TYPE: (CLINICAL TRIAL)

(CLINICAL TRIAL, PHASE I)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199709

ENTRY DATE: Entered STN: 16 Sep 1997

Last Updated on STN: 16 Sep 1997

Entered Medline: 2 Sep 1997

AB This was a phase I dose-finding and pharmacokinetic study of vinorelbine (Navelbine) and docetaxel (Taxotere) as first-line chemotherapy for metastatic breast cancer. Vinorelbine dose, 20 or 22.5 mg/m², on days 1 and 5, was followed on day 1 by docetaxel every 21 days, in doses increasing from 60 to 100 mg/m². Two maximum tolerated doses were reached, the first at 75 mg/m² of docetaxel and 22.5 mg/m² of vinorelbine, and the second at 100 mg/m² of docetaxel and 20 mg/m² of vinorelbine. Symptomatic peripheral neuropathy was not observed.

The recommended doses for phase II studies are 75 to 85 mg/m² of docetaxel on day 1 and 20 mg/m² of vinorelbine on days 1 and 5, every 3 weeks. The treatment regimen, which included 3-day corticosteroid prophylaxis, resulted in only mild fluid retention. Responses were seen at all dose levels, with an 80% overall response rate at the higher recommended dose; the overall response rate for patients at all dose levels was 66%. A high rate of response, including a complete response, was observed in patients with liver metastases.

L12 ANSWER 36 OF 55 MEDLINE on STN
ACCESSION NUMBER: 96150144 MEDLINE
DOCUMENT NUMBER: PubMed ID: 8546908
TITLE: A late phase II study of RP56976 (docetaxel) in patients with advanced or recurrent breast cancer.
AUTHOR: Adachi I; Watanabe T; Takashima S; Narabayashi M; Horikoshi N; Aoyama H; Taguchi T
CORPORATE SOURCE: Department of Medical Oncology, National Cancer Center Hospital, Tokyo, Japan.
SOURCE: British journal of cancer, (1996 Jan) Vol. 73, No. 2, pp. 210-6.
JOURNAL CODE: 0370635. ISSN: 0007-0920.
PUB. COUNTRY: SCOTLAND: United Kingdom
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199602
ENTRY DATE: Entered STN: 6 Mar 1996
Last Updated on STN: 6 Feb 1998
Entered Medline: 16 Feb 1996
AB A late phase II clinical trial of RP56976 (docetaxel), derived from Taxus baccata was performed to evaluate anti-tumour activity, time to progression and clinical toxicity in patients with advanced or recurrent breast cancer. The patients, between 15 and 80 years old with performance status (PS) of 0-2, received at least two cycles of docetaxel 60 mg m⁻² intravenously at 3-4 week intervals. Of the 81 patients enrolled, the 72 eligible for the study were given a total of 327 cycles, with a median of four cycles each. Five patients obtained a complete response (CR) and 27 a partial response (PR); the response rate (RR) was 44.4% (95% confidence interval 32.7-56.6%). A relatively high RR of 9/28 (32.1%) was observed in patients who had received prior chemotherapy involving anthracyclines. The dose-limiting toxicity was grade 3-4 leucocytopenia or neutropenia, found in 78.9% and 85.9% patients respectively. Other severe (grade > 3) toxicities included alopecia (38%), anorexia (18.3%), nausea/vomiting (11.3%), and fatigue (9.9%). Hypersensitivity reactions, oedema and skin toxicity were not severe and were reversible. One therapy-related death occurred 10 days after the initial dose was given. These findings indicate that docetaxel has potent activity against metastatic breast cancer, and that the dose of 60 mg m⁻² is safe.

L12 ANSWER 37 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2002:123598 CAPLUS.
DOCUMENT NUMBER: 136:161350
TITLE: Method of inhibiting angiogenesis associated with malignant and neoplastic cells using active vitamin D analogs
INVENTOR(S): Bishop, Charles W.; Mazess, Richard B.
PATENT ASSIGNEE(S): Bone Care International, Inc., USA
SOURCE: U.S. Pat. Appl. Publ., 13 pp., Cont.-in-part of U.S. Ser. No. 596,149.
CODEN: USXXCO
DOCUMENT TYPE: Patent
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 20

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-----------------|--------------|
| US 2002019375 | A1 | 20020214 | US 2001-891805 | 20010626 |
| US 6573256 | B2 | 20030603 | | |
| US 5763429 | A | 19980609 | US 1996-781910 | 19961230 <-- |
| US 6537982 | B1 | 20030325 | US 1998-596149 | 19980223 |
| PRIORITY APPLN. INFO.: | | | US 1996-781910 | A3 19961230 |
| | | | US 1998-596149 | A2 19980223 |
| | | | US 1993-119895 | A2 19930910 |
| | | | US 1994-265438 | A2 19940624 |
| | | | US 1995-415488 | A2 19950403 |
| | | | US 1995-486387 | A2 19950607 |

OTHER SOURCE(S): MARPAT 136:161350

AB Methods are disclosed which use active vitamin D analogs for the inhibition of angiogenesis associated with malignant and neoplastic cells. Methods comprise the application of an effective amount of a hypocalcemic hydroxyvitamin D compound to inhibit the angiogenesis of malignant cells, induce the apoptosis of malignant cells, and regress the growth of tumor cells.

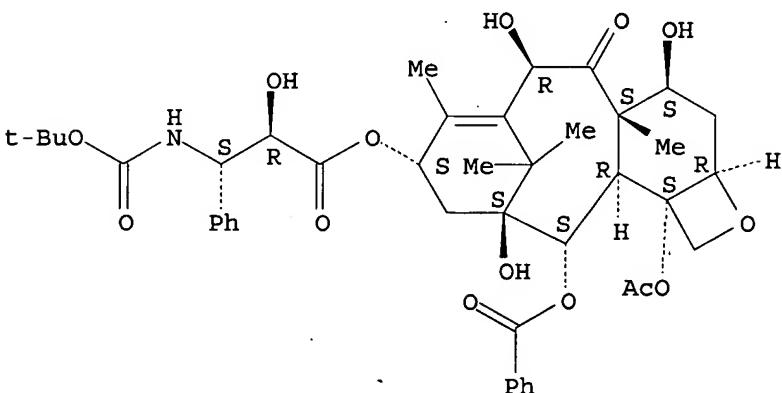
IT 114977-28-5, Docetaxel

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(hydroxyvitamin D compds. for inhibition of tumor-associated angiogenesis, and use with other agents)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 38 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:72803 CAPLUS

DOCUMENT NUMBER: 136:113175

TITLE: Method of treating malignancy-associated hypercalcemia using active vitamin D analogs

INVENTOR(S): Bishop, Charles W.; Mazess, Richard B.

PATENT ASSIGNEE(S): Bone Care International, Inc., USA

SOURCE: U.S. Pat. Appl. Publ., 12 pp., Cont.-in-part of U.S. 5,763,429.

CODEN: USXXCO

DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 20
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|--|-----------------|-----------------|--------------|
| US 2002010165 | A1 | 20020124 | US 2001-891763 | 20010626 |
| US 6566353 | B2 | 20030520 | | |
| US 5763429 | A | 19980609 | US 1996-781910 | 19961230 <-- |
| CA 2451037 | AA | 20030109 | CA 2002-2451037 | 20020626 |
| WO 2003002060 | A2 | 20030109 | WO 2002-US20320 | 20020626 |
| WO 2003002060 | A3 | 20040311 | | |
| W: | AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZW | | | |
| RW: | GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | |
| EP 1416939 | A2 | 20040512 | EP 2002-747979 | 20020626 |
| R: | AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | |
| CN 1520301 | A | 20040811 | CN 2002-812871 | 20020626 |
| JP 2004535439 | T2 | 20041125 | JP 2003-508302 | 20020626 |
| US 2003207810 | A1 | 20031106 | US 2003-441731 | 20030520 |
| PRIORITY APPLN. INFO.: | | | | |
| | | US 1996-781910 | A2 | 19961230 |
| | | US 1993-119895 | A2 | 19930910 |
| | | US 1994-265438 | A2 | 19940624 |
| | | US 1995-415488 | A2 | 19950403 |
| | | US 1995-486387 | A2 | 19950607 |
| | | US 1998-596149 | A3 | 19980223 |
| | | US 2001-891763 | A | 20010626 |
| | | WO 2002-US20320 | W | 20020626 |

OTHER SOURCE(S): MARPAT 136:113175

AB Methods utilizing active vitamin D analogs for the treatment of malignancy-associated hypercalcemia. Methods comprise the application of an effective amount of a hypocalcemic vitamin D compound to alleviate hypercalcemia, lower serum parathyroid hormone related protein (PTHrP) levels. The hypocalcemic vitamin D compds. can be coadministered with a cytotoxic agent.

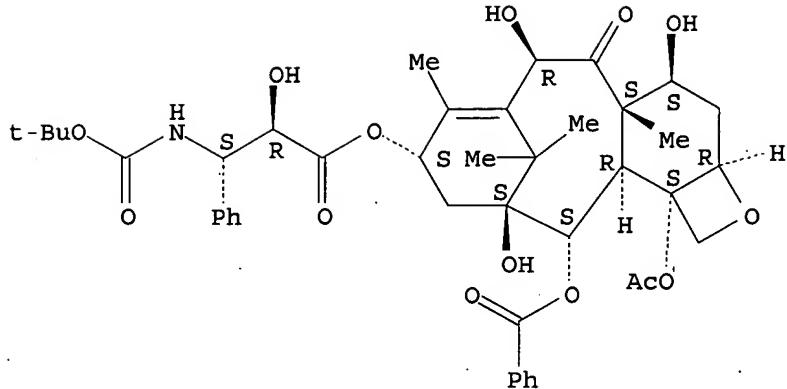
IT 114977-28-5, Docetaxel

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (method of treating malignancy-associated hypercalcemia using active vitamin D analogs coadministered with cytotoxic agents)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 39 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:507531 CAPLUS

DOCUMENT NUMBER: 135:107247

TITLE: Preparation of 3-heteroarylidenyl-2-indolinone compounds for modulating protein kinase activity and for use in cancer chemotherapy

INVENTOR(S): Langecker, Peter J.; Shawver, Laura K.; Tang, Peng C.; Sun, Li

PATENT ASSIGNEE(S): Sugen, Inc., USA

SOURCE: PCT Int. Appl., 93 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 2

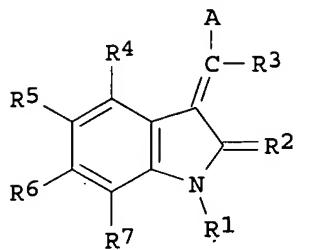
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|--|--|
| WO 2001049287 | A1 | 20010712 | WO 2000-US18058 | 20000630 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| WO 2000038519 | A1 | 20000706 | WO 1999-US31232 | 19991230 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| US 2003073837 | A1 | 20030417 | US 1999-476232 | 19991230 |
| EP 1259234 | A1 | 20021127 | EP 2000-943334 | 20000630 |
| EP 1259234 | B1 | 20060816 | | |
| R: AT, BE, CH, DE, DK, ES, FR, IE, SI, LT, LV, FI, RO, MK, CY, AL | | | | |
| JP 2003535038 | T2 | 20031125 | JP 2001-549655 | 20000630 |
| US 2003191162 | A1 | 20031009 | US 2002-307483
US 1999-476232
WO 1999-US31232
US 2000-569545
US 1998-114313P | 20021202
A 19991230
A 19991230
A 20000512
P 19981231 |
| PRIORITY APPLN. INFO.: | | | | |

OTHER SOURCE(S) :
GI

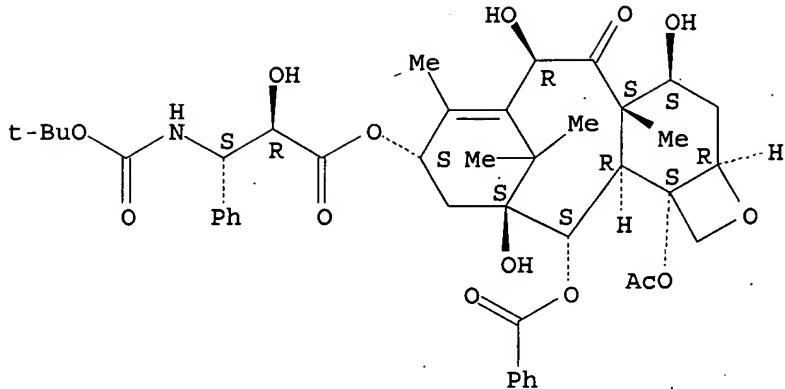
MARPAT 135:107247

WO 2000-US18058 W 20000630



- AB The present invention relates to 3-heteroarylidenyl-2-indolinone compds. [I; R1 = H, alkyl; R2 = O, S; R3 = H; R4 , R5, R6, R7 = H, alkyl, alkoxy, aryl, aryloxy, alkaryloxy, halo, trihalomethyl, S(O)R, SO2NRR', SO3R, SR, NO2, NRR', OH, cyano, COR, O2CR, (CH2)nCO2R, CONRR'; A = a five membered heteroaryl selected from (un)substituted thiophene, pyrrole, pyrazole, imidazole, 1,2,3-triazole, 1,2,4-triazole, oxazole, isoxazole, thiazole, isothiazole, 2-sulfonylfuran, 4-alkylfuran, 1,2,3-oxadiazole, 1,2,4-oxadiazole, 1,2,5-oxadiazole, 1,3,4-oxadiazole, 1,2,3,4-oxatriazole, 1,2,3,5-oxatriazole, 1,2,3-thiadiazole, etc.; n = 0-3; R, R' = H, alkyl, aryl] or physiol. acceptable salts or prodrugs thereof are prepared. These compds. modulate the enzymic activity of protein kinases such as receptor protein tyrosine kinase, cellular tyrosine kinase, and serine threonine kinase and therefore are expected to be useful in the prevention and treatment of protein kinase related cellular disorders such as cancer. Furthermore, these compds. are expected to enhance the efficacy of other chemotherapeutic agents, in particular, fluorinated pyrimidines, in the treatment of cancer. In a cellular-based assay for inhibiting the receptor phosphorylation, 3-[(2,4-dimethylpyrrol-5-yl)methylidenyl]-2-indolinone (II) inhibited Flk-1-autophosphorylation with IC50 of .apprx.1 μ M. II in vitro inhibited proliferation of endothelial cells induced by VEGF with IC50 of .apprx.0.07 μ M. Although II in vitro had no direct inhibitory effect on a variety of tumor cell lines at concentration up to
- 50 μ M, it in vivo demonstrated a significant suppression of tumor growth against a broad spectrum of tumor types s.c. implanted into immunocompromised mice and whose growth are driven by various growth factors such as PDGF, EGF, and Her2.
- IT 114977-28-5, Docetaxel
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(for cancer chemotherapy in combination with heteroarylidenylindolinone derivative; preparation of 3-heteroarylidenyl-2-indolinone compds. for modulating protein kinase activity for cancer chemotherapy)
- RN 114977-28-5 CAPLUS
- CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 40 OF 55 MEDLINE on STN

ACCESSION NUMBER: 2000024226 MEDLINE

DOCUMENT NUMBER: PubMed ID: 10560436

TITLE: A case of hepatic arterial infusion chemotherapy with docetaxel for liver metastasis from breast cancer.

AUTHOR: Kim S J; Maeura Y; Ueda N; Saito M; Matsunaga S

CORPORATE SOURCE: Senri Hoken Medical Center, Dept. of Surgery, Shinsenri Hospital.

SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1999 Oct) Vol. 26, No. 12, pp. 1959-62.

Journal code: 7810034. ISSN: 0385-0684.

PUB. COUNTRY: Japan

DOCUMENT TYPE: (CASE REPORTS)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Japanese

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199911

ENTRY DATE: Entered STN: 13 Jan 2000

Last Updated on STN: 13 Jan 2000

Entered Medline: 26 Nov 1999

AB We experienced a case of hepatic arterial infusion chemotherapy using docetaxel for liver metastasis, which showed no response to CEF therapy, from breast cancer. A 63-year-old woman had undergone modified radical mastectomy for right breast cancer (T2aN1bM0: Stage II) in October, 1995. Six-cycle CMF therapy and toremifene citrate (40 mg/day) were administered as adjuvant therapy, but multiple recurrent tumors in liver, lung, and local site were detected in February 1997. Six-cycle CEF therapy was given for recurrent disease and there was a complete response for lung and local recurrence, but no change in liver metastasis. Chemoendocrine therapies using 5'-DFUR or CMitF in addition to TAM and fadrozole hydrochloride hydrate had developed progressive disease for liver metastasis. A catheter and port kit were operatively inserted and implanted in March 1998. Hepatic arterial infusion of docetaxel (30-40 mg/body/month, one hour administration) was repeated 4 times, once in our clinic. Leukopenia, general fatigue and fever, which were mild and did not require any treatment, appeared as side effects. This treatment reduced multiple liver metastatic sites on abdominal CT finding and was thought to be a partial response. However, the patient had multiple brain metastasis and died on August 2, 1998. While docetaxel, even by systemic administration, has a 36-77% response rate for liver metastasis, arterial infusion might have a good response and mild side effect with a lower dose than by intravenous administration.

L12 ANSWER 41 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:161407 CAPLUS

DOCUMENT NUMBER: 134:202681
 TITLE: Dietary supplementation with, and methods for,
 administration of a yeast-derived selenium product,
 and use in cancer chemotherapy
 INVENTOR(S): Hsia, Houn Simon; Yang, Ping; Arnold, Michael
 PATENT ASSIGNEE(S): Viva America Marketing Corporation, USA
 SOURCE: U.S., 9 pp., Cont.-in-part of U.S. 6,140,107.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 4
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-----------------|--------------|
| US 6197295 | B1 | 20010306 | US 1999-303993 | 19990503 |
| US 6140107 | A | 20001031 | US 1996-719572 | 19960925 <-- |
| US 6368643 | B1 | 20020409 | US 1999-298114 | 19990423 |
| US 2001043925 | A1 | 20011122 | US 2001-801124 | 20010305 |
| US 6576233 | B2 | 20030610 | | |
| PRIORITY APPLN. INFO.: | | | US 1996-719572 | A2 19960925 |
| | | | US 1997-802773 | B2 19970221 |
| | | | US 1998-15758 | A2 19980129 |
| | | | US 1998-82939P | P 19980424 |
| | | | US 1999-303993 | A3 19990503 |

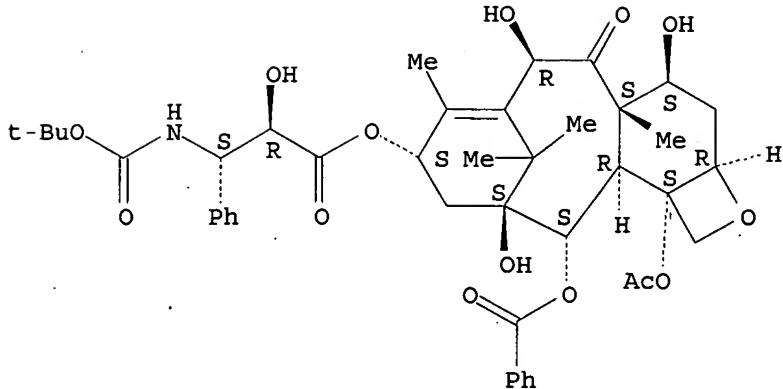
AB The invention solves the need for nontoxic forms of selenium which is an essential part of the human diet. The invention provides dried-yeast products containing selenium, as well as a method of producing the dried yeast products. The method uses selenium having high biol. activity but low toxicity. The invention also provides nutritional supplements containing the selenium-containing dried yeast products and methods of administering these products and supplements to improve human health. The invention also provides a practically nontoxic yeast selenium product having increased intracellular selenium concns., as well as methods to reduce tumor cell growth by administration of a selenium yeast product comprising yeast *Saccharomyces boulardii sequela PY31 (ATCC 74366)* in combination with chemotherapeutic agents.

IT 114977-28-5, Taxotere
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (dietary supplementation with yeast-derived selenium product, and use in cancer chemotherapy)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy) carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 42 OF 55 MEDLINE on STN

ACCESSION NUMBER: 2001191709 MEDLINE

DOCUMENT NUMBER: PubMed ID: 11124653

TITLE: Chemotherapy-induced noncardiogenic pulmonary edema related to gemcitabine plus docetaxel combination with granulocyte colony-stimulating factor support.

AUTHOR: Briassoulis E; Froudarakis M; Milionis H J; Peponis I; Constantopoulos S; Pavlidis N

CORPORATE SOURCE: Department of Medical Oncology, Ioannina University Hospital, Ioannina, Greece.. ebriasou@otenet.gr

SOURCE: Respiration; international review of thoracic diseases, (2000) Vol. 67, No. 6, pp. 680-3.
Journal code: 0137356. ISSN: 0025-7931.

PUB. COUNTRY: Switzerland

DOCUMENT TYPE: (CASE REPORTS)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200104

ENTRY DATE: Entered STN: 10 Apr 2001

Last Updated on STN: 10 Apr 2001

Entered Medline: 5 Apr 2001

AB Several cancer therapeutic agents have been associated with pulmonary toxicity. Herein, we describe the case of a 73-year-old woman with breast cancer metastatic to the liver, who developed noncardiogenic pulmonary edema (NPE) while on treatment with gemcitabine plus docetaxel combination with granulocyte colony-stimulating factor (G-CSF) support. Gemcitabine, a deoxycytidine analogue, is reported to produce mild self-limiting and only occasionally severe pulmonary toxicity. The microtubule stabilizer docetaxel has been associated with water retention complications. The combination of these two agents has shown promising activity in several solid tumors and is in a phase of clinical development with prophylactic G-CSF in most of the trials due to the high rate of dose-limiting neutropenia observed with this combination. In our case pulmonary toxicity resolved rapidly following the administration of corticosteroids. A possible deleterious synergy of the compounds involved in this case is discussed and the medical literature on NPE related to cancer therapy is shortly reviewed. We conclude that NPE should always be considered in patients with respiratory function deterioration while on therapy with the gemcitabine-docetaxel combination and G-CSF. Corticosteroids can provide maximum benefit if started early upon diagnosis coupled with withdrawal of the causative drugs.

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L12 ANSWER 43 OF 55 MEDLINE on STN

ACCESSION NUMBER: 1999197818 MEDLINE
DOCUMENT NUMBER: PubMed ID: 10097745
TITLE: A late phase II clinical study of RP56976 (docetaxel) in patients with advanced or recurrent gastric cancer: a cooperative study group trial (group B).
AUTHOR: Mai M; Sakata Y; Kanamaru R; Kurihara M; Suminaga M; Ota J; Hirabayashi N; Taguchi T; Furue H
CORPORATE SOURCE: Dept. of Surgery, Cancer Research Institute, Kanazawa University.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1999 Mar) Vol. 26, No. 4, pp. 487-96.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
(MULTICENTER STUDY)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199904
ENTRY DATE: Entered STN: 13 Apr 1999
Last Updated on STN: 13 Apr 1999
Entered Medline: 1 Apr 1999
AB A late phase II clinical study of RP56976 (docetaxel) in patients with advanced or recurrent gastric cancer was performed to evaluate the anti-tumor activity and clinical toxicity as a multicenter cooperative trial. Docetaxel was administered intravenously at a dose of 60 mg/m² every 3-4 weeks. Of 72 patients enrolled, 63 patients were eligible and 59 patients were evaluable for response. The anti-tumor effects obtained complete response (CR) in one patient partial response (PR) in 13, minor response (MR) in 3, no change (NC) in 20, and progressing disease (PD) in 22 patients. The overall response rate in 59 patients was 23.7% (14/59). For 14 CR or PR cases, a response appeared 10 to 107 days (median 33.5 days) and 1 to 8 (median 2) times of dosing after the initial administration. The response rate was 9.5% in the primary tumor, 31.3% livers, 50.0% abdominal tumor, and 24.1% lymph nodes, respectively. The major adverse reactions were gastrointestinal symptoms including nausea/vomiting, anorexia, fatigue, alopecia and fever. Leukocytopenia and neutrocytopenia were also observed with a high incidence, but they recovered after 8 days from the nadir. The results show that docetaxel is an effective anti-tumor agent for advanced or recurrent gastric cancer. It is necessary to conduct another clinical trial by concomitant administration with other anti-tumor agents.

L12 ANSWER 44 OF 55 MEDLINE on STN
ACCESSION NUMBER: 1999014548 MEDLINE
DOCUMENT NUMBER: PubMed ID: 9797814
TITLE: Late phase II clinical study of RP56976 (docetaxel) in patients with advanced/recurrent gastric cancer: a Japanese Cooperative Study Group trial (group A).
AUTHOR: Taguchi T; Sakata Y; Kanamaru R; Kurihara M; Suminaga M; Ota J; Hirabayashi N
CORPORATE SOURCE: Japan Society for Cancer Chemotherapy, Aomori Prefectural Central Hospital.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1998 Oct) Vol. 25, No. 12, pp. 1915-24.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
(MULTICENTER STUDY)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals

ENTRY MONTH: 199811
ENTRY DATE: Entered STN: 6 Jan 1999
Last Updated on STN: 6 Jan 1999
Entered Medline: 4 Nov 1998

AB A late phase II clinical study of RP56976 (docetaxel) was conducted in patients with advanced/recurrent gastric cancer as a multicenter cooperative trial. Docetaxel was administered intravenously at a dose of 60 mg/m² every 3-4 weeks. Of the 76 patients enrolled, 66 patients were eligible and 59 patients were evaluable for response. One patient showed complete response (CR), 13 patients partial response (PR), 1 patient minor response (MR), 19 patients no change (NC) and 25 patients had progressive disease (PD). The overall response rate in 59 evaluable patients was 23.7% (95% CI = 13.6-36.6%). The primary tumor showed a 4.3% (1/23) response, while the metastatic lesions in the abdomen, pelvic mass, lung, liver, and lymph nodes showed response rates of 62.5% (5/8), 33.3% (1/3), 33.3% (1/3), 14.8% (4/27), and 13.9% (5/26), respectively. About hematological toxicity, severe (Grade 3 or more) leukopenia was observed in 36 patients (56.3%) and neutropenia in 52 patients (81.3%). Other major toxicity (Grade 3 or more) included nausea/vomiting in 11 patients (17.2%), anorexia in 9 patients (14.1%), fatigue in 5 patients (7.8%), and alopecia in 7 patients (10.9%), all which were tolerable. The results show that docetaxel is an effective anticancer agent for advanced/recurrent gastric cancer.

L12 ANSWER 45 OF 55 MEDLINE on STN
ACCESSION NUMBER: 2000464139 MEDLINE
DOCUMENT NUMBER: PubMed ID: 11016005
TITLE: A case of effective chemotherapy using CAF followed by docetaxel for advanced breast cancer.
AUTHOR: Kokufu I; Taniguchi H; Kim Y H; Fukuda K; Yamamoto M; Yano T; Yamada K; Kitano H; Fukuda H
CORPORATE SOURCE: Dept. of Surgery, Itami City Hospital.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (2000 Sep) Vol. 27, No. 10, pp. 1577-80.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CASE REPORTS)
Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200010
ENTRY DATE: Entered STN: 19 Oct 2000
Last Updated on STN: 19 Oct 2000
Entered Medline: 10 Oct 2000

AB A huge mass measuring 13 x 12 cm and wide cutaneous edema were detected in the right breast of a 51-year-old woman. Under a diagnosis of locally advanced breast cancer (T4bN2M1, stage IV) with liver metastases, we attempted sequential neoadjuvant chemotherapy. After three courses of CAF therapy (cyclophosphamide, doxorubicin (DXR), 5-FU), the primary tumor was decreased by 56% and the liver metastases had disappeared. A minor pathologic response was observed. Subsequently, three courses of docetaxel (TXT) administration were carried out. The primary tumor was then decreased by 75% and the axillary metastases had disappeared. Histopathological examination showed gross viable tumor cells in the residual tumor and positive axillary lymph nodes. The only toxic effect was nausea (grade 1) and no major adverse effects were observed. Neoadjuvant chemotherapy with sequential DXR followed by TXT is a useful treatment for locally advanced breast cancer.

L12 ANSWER 46 OF 55 MEDLINE on STN
ACCESSION NUMBER: 1999430325 MEDLINE
DOCUMENT NUMBER: PubMed ID: 10500538
TITLE: A case of recurrent breast cancer successfully treated with docetaxel.

AUTHOR: Koshizuka K; Hada M; Muto S; Hagiwara J; Nakagomi H; Takano K; Kamiya K; Tada Y
CORPORATE SOURCE: Second Dept. of Surgery, Yamanashi Medical University.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1999 Sep) Vol. 26, No. 10, pp. 1479-81.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CASE REPORTS)
Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199909
ENTRY DATE: Entered STN: 12 Oct 1999
Last Updated on STN: 12 Oct 1999
Entered Medline: 30 Sep 1999

AB A 53-year-old female underwent mastectomy for left breast cancer in April, 1993. She was given oral tamoxifen but this had to be discontinued due to its side effects. In March, 1998, she developed bone and lung metastases, in spite of treatment with combination chemotherapy (CEF). We thus treated here with docetaxel 90 mg three times and 40 mg six times. After the chemotherapy, she achieved complete remissions of the lung metastases and a decrease in serum CEA, CA 15-3, NCC-ST439, and BCA225. Adverse reactions to docetaxel were grade 2 alopecia, grade 4 neutropenia, dysgeusia, and fluid retention. All were tolerable.. This new agent may play an important future role in chemotherapy for recurrent breast cancer.

L12 ANSWER 47 OF 55 MEDLINE on STN
ACCESSION NUMBER: 1998233482 MEDLINE
DOCUMENT NUMBER: PubMed ID: 9571974
TITLE: Breast cancer with liver metastasis responsive to docetaxel: case report.
AUTHOR: Oura S; Sakurai T; Yoshimura G; Tamaki T; Umemura T; Kokawa Y
CORPORATE SOURCE: Dept. of Surgery, Wakayama Medical College Kihoku Hospital.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1998 Apr) Vol. 25, No. 5, pp. 743-6.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CASE REPORTS)
Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199805
ENTRY DATE: Entered STN: 14 May 1998
Last Updated on STN: 14 May 1998
Entered Medline: 7 May 1998

AB A 59-year-old female underwent mastectomy for right breast cancer in November 1992. She received tamoxifen and anthracycline-containing chemotherapy as adjuvant therapy. In and after September 1994, she developed loco-regional recurrences five times in total, each of which was treated with surgery and conventional combination chemotherapy. In April 1997, she developed liver metastasis, which was refractory to biochemical modulation therapy (low-dose cisplatin + 5-FU). We, therefore, treated her six times with docetaxel 80 mg, which resulted in partial response of the liver metastasis and brought about a marked decrease in serum CA15-3 levels. Adverse effects of docetaxel were grade 3 alopecia and leucocytopenia. She has been well without re-growth of the liver metastasis for over five months.

L12 ANSWER 48 OF 55 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
ACCESSION NUMBER: 1998:259786 BIOSIS
DOCUMENT NUMBER: PREV199800259786

TITLE: Antitumour effect of docetaxel in malignant diseases.
 AUTHOR(S): Eckhardt, Sandor [Reprint author]
 CORPORATE SOURCE: Rath Gyorgy u. 7-9, 1122 Budapest, Hungary
 SOURCE: Orvosi Hetilap, (April 12, 1998) Vol. 139, No. 15, pp. 867-872. print.
 CODEN: ORHEAG. ISSN: 0030-6002.
 DOCUMENT TYPE: Article
 General Review; (Literature Review)
 LANGUAGE: Hungarian
 ENTRY DATE: Entered STN: 9 Jun 1998
 Last Updated on STN: 12 Aug 1998
 AB In recent years numerous molecular biological discoveries enlightened the various steps of the neoplastic transformation. Based on new targets, this development made it possible to synthetize new tumour inhibitory substances. Among them taxanes capable to block depolymerization of tubulin - which is an essential molecule in cell division - play an important role. Docetaxel (Taxotere) belongs to this group and is an active drug in the treatment of breast cancer. Moreover, platinum-resistant tumours may also respond to the therapy. It is important to note that even visceral (hepatic) metastases may express chemosensitivity. Results of combination chemotherapy seem to be also promising. The antitumour effect of Taxotere in NSCLC and other malignant neoplasms is under investigation. The toxicity of Taxotere may be successfully reduced by premedication of steroids. The necessary protective measures render the Taxotere therapy safe and of being perspectivistic.

L12 ANSWER 49 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1999:325975 CAPLUS
 DOCUMENT NUMBER: 130:357177
 TITLE: Detoxication of active pharmaceutical substances using cyclodextrin oligomers
 INVENTOR(S): Moser, Joerg G.
 PATENT ASSIGNEE(S): Germany
 SOURCE: PCT Int. Appl., 31 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: German
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------------------|-------------|
| WO 9924474 | A1 | 19990520 | WO 1998-EP7229 | 19981111 <- |
| W: AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, HR, HU, ID, IL, IS, JP, KP, KR, LC, LK, LR, LS, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 9916694 | A1 | 19990531 | AU 1999-16694 | 19981111 <- |
| EP 1045863 | A1 | 20001025 | EP 1998-961184 | 19981111 <- |
| EP 1045863 | B1 | 20030402 | | |
| R: AT, BE, CH, DE, ES, FR, GB, IT, LI, NL | | | | |
| JP 2001522901 | T2 | 20011120 | JP 2000-520482 | 19981111 |
| AT 236195 | E | 20030415 | AT 1998-961184 | 19981111 |
| US 6642214 | B1 | 20031104 | US 2000-554223 | 20000803 |
| PRIORITY APPLN. INFO.: | | | DE 1997-19749801 A 19971111 | |
| | | | DE 1998-19822416 A 19980519 | |
| | | | WO 1998-EP7229 W 19981111 | |

AB Cyclodextrin oligomers with 2 cyclodextrins connected via a spacer B on the secondary side [CD-X-A-X-B-X-A-X-CD; CD = cyclodextrin; X = bond, NH,

O, S, C(O); A = bond, C2-4 aliphatic residue; B = rigid, preferably hydrophilic residue] form strongly hydrophilic inclusion compds. with pharmaceutical agents and thereby prevent toxic side effects of drugs on nontarget cells by inhibiting their uptake into the cells. The drugs can be targeted to specific tissue sites by attachment of affinity groups such as antibodies to the cyclodextrin residues, and the drug can be released at the target site by destruction of the cyclodextrin residues (e.g. with cyclodextrinase from Klebsiella oxytoca). Provided the cyclodextrins are connected on their secondary sides, their cavities will face each other; the distance between them is determined by the choice of spacer, and is preferably 0.8-1.8 nm. Thus, β -cyclodextrin was condensed with 4,4'-methylenebis(benzenesulfonyl chloride) and the product reacted with diaminopropane to form β -6(A-D)-diamidopropanediaminocyclodextrin (I). Sep., 2-monotosyl- β -cyclodextrin reacted with 3-mercaptopropionic acid to form β -(2)cyclodextrin-(3-thiopropionic acid) (II). Reaction of II with carbonyldimidazole, N-hydroxysuccinimide, and a 2.5-fold molar excess of I produced a cyclodextrin trimer. Nude mice bearing OAT SCLC cell tumors were treated with biotinylated monoclonal antibody ICO 25 i.p., followed 24 h later by NeutrAvidin i.p., and after an addnl. 48 h by a biotinylcadaverine-labeled CD dimer-paclitaxel complex. Growth of the tumors was inhibited without occurrence of side effects.

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 50 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1998:213711 CAPLUS

DOCUMENT NUMBER: 128:289570

TITLE: Pharmacokinetics of anticancer agents in patients with impaired liver function

AUTHOR(S): Donelli, M. G.; Zucchetti, M.; Munzone, E.; D'incalci, M.; Crosignani, A.

CORPORATE SOURCE: Dipartimento di Oncologia, Istituto di Ricerche Farmacologiche Mario Negri, Milan, 20157, Italy

SOURCE: European Journal of Cancer (1998), 34(1), 33-46

PUBLISHER: CODEN: EJCAEL; ISSN: 0959-8049
Elsevier Science Ltd.

DOCUMENT TYPE: Journal; General Review

LANGUAGE: English

AB A review with 95 refs. This report reviews published information on the clin. pharmacokinetics of antitumor agents in patients with liver dysfunction, associated with primary liver disease or liver metastases. Information was available for anthracyclines and their related compds., antimetabolites, cyclophosphamide, vinca alkaloids, taxanes and epipodophyllotoxins. Changes in the pharmacokinetic profile or metabolism in patients with mild or severe hepatobiliary dysfunction are described and the relationships between serum levels, parameters employed for measuring hepatic function and toxic or therapeutic effects are examined. Current knowledge of the pharmacokinetics of antineoplastic agents in liver disease is far from complete, mostly obtained in small nos. of non-homogeneous patients often presenting only moderate liver dysfunction, and empirical guidelines for dose assessment are still largely applied in clin. practice. Because of the complex pathophysiol. mechanisms of liver insufficiency in cancer patients, there is still doubt whether endogenous markers are useful. Although caution in treating cancer patients with liver insufficiency is compulsory, for most compds. there seems no need to recommend dose redns. for moderate impairment. However, for the tubulin acting agents, vincristine, vinblastine and possibly for paclitaxel and docetaxel, there is strong evidence that dose adjustment is mandatory in order to avoid excessive neutropenia and neurotoxicity.

REFERENCE COUNT: 95 THERE ARE 95 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 51 OF 55 MEDLINE on STN
ACCESSION NUMBER: 2000390579 MEDLINE
DOCUMENT NUMBER: PubMed ID: 10895201
TITLE: Preliminary results of multicenter phase II trial of docetaxel (Taxotere) in combination with doxorubicin as first line chemotherapy in Indonesian patients with advanced or metastatic breast cancer.
AUTHOR: Muthalib A; Darwis I; Prayogo N; Sutjipto
CORPORATE SOURCE: Dharmais National Cancer Center/School of Medicine, University of Indonesia, Jakarta.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (2000 May) Vol. 27 Suppl 2, pp. 498-504.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
(MULTICENTER STUDY)
(RANDOMIZED CONTROLLED TRIAL)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200008
ENTRY DATE: Entered STN: 18 Aug 2000
Last Updated on STN: 18 Aug 2000
Entered Medline: 10 Aug 2000
AB RATIONALE: Docetaxel and doxorubicin have produced a high degree of activity in previously untreated/treated patients with metastatic breast cancer (MBC). The efficacy of Taxotere (T) single agent as 2nd line chemotherapy is well established in large randomized phase III studies. OBJECTIVE: The objective of this study is to confirm the efficacy and safety of a combination of Taxotere with doxorubicin as 1st line chemotherapy in Indonesian MBC patients. TREATMENT AND METHOD: Eighteen patients age < or = 70 years with advanced or metastatic breast cancer (MBC) with no prior taxane chemotherapy or prior cumulative doxorubicin (D) of no more than 250 mg/m² and no heart disease were enrolled in this phase II study of D (50 mg/m²) IV bolus followed one hour later by Taxotere (T) 60 mg/m² IV infusion over 1 hour every 3 weeks for 6 cycles treatments. A 3-day oral corticosteroid premedication was administered starting one day before the infusion of each cycle. Left ventricular ejection fraction (LVEF) was evaluated at baseline and after cycle 6. PATIENTS CHARACTERISTICS: 18 patients (pts) have been treated with 108 cycles administered. Median age was 46 years (31-58), WHO PS 0 = 50%, 1 = 50% and number of organs involved were: 2 (72%), 3 (22%) and 4 (6%). RESULTS: After 3 cycles, partial (PR) and no change (NC) responses occurred in 15 pts (83.3%) and 3 pts (16.7%). The best overall response after 6 cycles, including complete (CR) and partial (PR) responses, occurred in 13 pts (72.2%) including 3 CRs and 10 PRs. Two patients with extensive liver metastases at the baseline had a complete disappearance after 6 cycles. No patients developed congestive heart failure (CHF). Grade 3/4 hematological toxicities included leukopenia in 18 pts (100%), febrile neutropenia in 6 pts (33%), leukopenia with infection in 2 pts (11%), leukopenia with fever in 1 pt (5.5%), and anemia in 6 pts (33.3%). Nonhematological toxicities grade 3/4 included alopecia (61%), asthenia (4.6%), nausea/vomiting (2.7%), pain (2.7%), stomatitis (2.7%), and diarrhoea (0.9%). Leukopenia was generally of short duration, occurred mainly during the first and second cycle, and did not require any dose reduction. There was one death due to progressive disease after six cycles of treatment. CONCLUSION: Taxotere-doxorubicin combination is very active in the first-line treatment of MBC, seems to be especially effective in patients with liver metastases, and is associated with a manageable toxicity profile.

ACCESSION NUMBER: 2001:89699 BIOSIS
DOCUMENT NUMBER: PREV200100089699
TITLE: Phase I study of weekly docetaxel in combination with capecitabine in patients with solid malignancies.
AUTHOR(S): Villalona-Calero, M. A. [Reprint author]; Shapiro, C. [Reprint author]; Otterson, G. A. [Reprint author]; Hauger, M. [Reprint author]; Kraut, E. [Reprint author]; Clinton, S. [Reprint author]; Shah, M. [Reprint author]; Stanek, M. [Reprint author]; Monk, J. P. [Reprint author]
CORPORATE SOURCE: Arthur James Cancer Center and R Solove Research Institute, Ohio State University, Columbus, OH, USA
SOURCE: Breast Cancer Research and Treatment, (November, 2000) Vol. 64, No. 1, pp. 125. print.
Meeting Info.: 23rd Annual San Antonio Breast Cancer Symposium. San Antonio, Texas, USA. December 06-09, 2000. Cancer Therapy and Research Center Research Foundation.
CODEN: BCTR6. ISSN: 0167-6806.
DOCUMENT TYPE: Conference; (Meeting)
Conference; Abstract; (Meeting Abstract)
LANGUAGE: English
ENTRY DATE: Entered STN: 14 Feb 2001
Last Updated on STN: 12 Feb 2002

L12 ANSWER 53 OF 55 MEDLINE on STN
ACCESSION NUMBER: 2000339901 MEDLINE
DOCUMENT NUMBER: PubMed ID: 10885392
TITLE: Metastasectomy as a cytoreductive strategy for treatment of isolated pulmonary and hepatic metastases from breast cancer.
AUTHOR: Bathe O F; Kaklamanos I G; Moffat F L; Boggs J; Franceschi D; Livingstone A S
CORPORATE SOURCE: Department of Surgery, University of Miami, FL 33136, USA.. bathe@worldnet.att.net
SOURCE: Surgical oncology, (1999 Jul) Vol. 8, No. 1, pp. 35-42. Ref: 45
Journal code: 9208188. ISSN: 0960-7404.
PUB. COUNTRY: Netherlands
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE).
General Review; (REVIEW)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200007
ENTRY DATE: Entered STN: 28 Jul 2000
Last Updated on STN: 28 Jul 2000
Entered Medline: 20 Jul 2000

AB The authors sought to examine the utility of resection in conjunction with adjuvant chemotherapy for treatment of metastases from breast cancer isolated to the liver or lungs. Limitations of regional therapy were examined and potential agents for systemic therapy were reviewed. As resection of metastases is a controversial therapeutic approach, no clinical trials are available for review. Rather, evidence for a potential role for surgery rests on retrospective studies of small series of patients. Technical advances have rendered resection of liver and lung metastases safe. Long-term results as reported by other investigators support the role of metastasectomy in selected patients. The site of failure following ablation of liver metastases is usually in the liver. Following resection of lung metastases, nonpulmonary and disseminated recurrences are most common. Adjuvant therapy with docetaxel or any other agent or combination with significant activity against visceral metastases might potentiate long-term results.

L12 ANSWER 54 OF 55 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
ACCESSION NUMBER: 2001:77815 BIOSIS

DOCUMENT NUMBER: PREV200100077815
TITLE: A phase II trial of escalated dose docetaxel (TXT) with G-CSF support in patients (pts) with advanced breast cancer.
AUTHOR(S): Mitchell, P. [Reprint author]; Bassar, R.; Harris, M. [Reprint author]; Ng, S.; Gibbs, P. [Reprint author]; Chipman, M. [Reprint author]; Grigg, A.; Jeffrey, A.; James, R.; Gargano, J.; Riva, A.; Appia, F.; Green, M.
CORPORATE SOURCE: Medical Oncology, Austin and Repatriation Medical Centre, Heidelberg West, VIC, Australia
SOURCE: Breast Cancer Research and Treatment, (November, 2000) Vol. 64, No. 1, pp. 88. print.
Meeting Info.: 23rd Annual San Antonio Breast Cancer Symposium. San Antonio, Texas, USA. December 06-09, 2000. Cancer Therapy and Research Center Research Foundation.
CODEN: BCTR D6. ISSN: 0167-6806.
DOCUMENT TYPE: Conference; (Meeting)
Conference; Abstract; (Meeting Abstract)
LANGUAGE: English
ENTRY DATE: Entered STN: 7 Feb 2001
Last Updated on STN: 12 Feb 2002

L12 ANSWER 55 OF 55 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
ACCESSION NUMBER: 2001:132253 BIOSIS
DOCUMENT NUMBER: PREV200100132253
TITLE: Close correlation of paraneoplastic hyperfibrinolysis with relapse and remission of anaplastic small cell carcinoma: A case report.
AUTHOR(S): Kegel, T. [Reprint author]; Kellner, O. [Reprint author]; Grothey, A. [Reprint author]; Wolf, H.-H. [Reprint author]; Voigt, W. [Reprint author]; Dorligshaw, O. [Reprint author]; Schmoll, H.-J. [Reprint author]
CORPORATE SOURCE: Dept. of Hematology/Oncology, University of Halle, Halle, Germany
SOURCE: Onkologie, (October, 2000) Vol. 23, No. Sonderheft 7, pp. 184. print.
Meeting Info.: Annual Meeting of the German and Austrian Society for Hematology and Oncology. Graz, Austria. October 21-25, 2000.
CODEN: ONKOD2. ISSN: 0378-584X.
DOCUMENT TYPE: Conference; (Meeting)
Conference; Abstract; (Meeting Abstract)
LANGUAGE: English
ENTRY DATE: Entered STN: 14 Mar 2001
Last Updated on STN: 15 Feb 2002

~~610 1000 P~~

L1 ANSWER 1 OF 1 REGISTRY COPYRIGHT 2006 ACS on STN

RN 114977-28-5 REGISTRY

ED Entered STN: 25 Jun 1988

CN Benzenepropanoic acid, β -[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN 7,11-Methano-1H-cyclodeca[3,4]benz[1,2-b]oxete, benzenepropanoic acid deriv.

CN Benzenepropanoic acid, β -[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, 12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, [2aR-[2a α ,4 β ,4a β ,6 β ,9 α (α R*, β S*)],11 α ,12 α ,12a α ,12b α]-

OTHER NAMES:

CN Docetaxel

CN Docetaxol

CN N-Debenzoyl-N-tert-butoxycarbonyl-10-deacetyltaxol

CN RP 56976

CN Taxotere

FS STEREOSEARCH

DR 216252-50-5

MF C43 H53 N O14

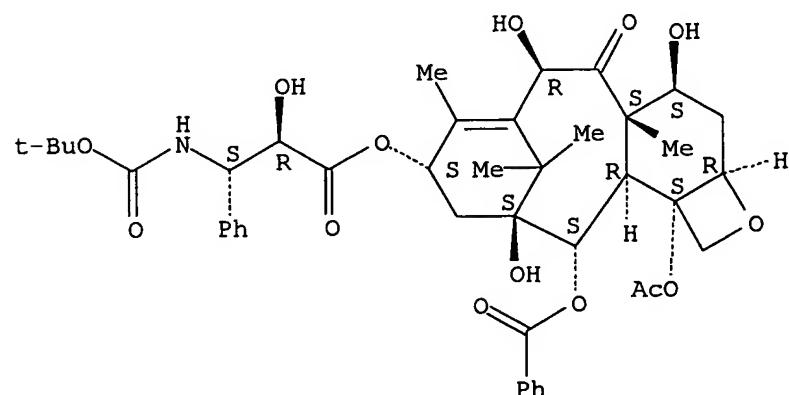
CI COM

SR CA

LC STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BEILSTEIN*, BIOSIS, BIOTECHNO, CA, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMINFORMRX, CIN, CSCHEM, DDFU, DRUGU, EMBASE, HSDB*, IMSCOSEARCH, IMSDRUGNEWS, IMSPATENTS, IMSRESEARCH, IPA, MEDLINE, MRCK*, MSDS-OHS, PATDPASPC, PHAR, PIRA, PROMT, PROUSDDR, PS, RTECS*, SYNTHLINE, TOXCENTER, USAN, USPAT2, USPATFULL

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Absolute stereochemistry.



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3480 REFERENCES IN FILE CAPLUS (1907 TO DATE)

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=> s docetaxol or docetaxel or txotere or rp 56976 or 114977-28-5
L1 20746 DOCETAXOL OR DOCETAXEL OR TXOTERE OR RP 56976 OR 114977-28-5

=> s docetaxol or docetaxel or txotere or rp 56976 or 114977-28-5/rn
'RN' IS NOT A VALID FIELD CODE
'RN' IS NOT A VALID FIELD CODE
'RN' IS NOT A VALID FIELD CODE
L2 18876 DOCETAXOL OR DOCETAXEL OR TXOTERE OR RP 56976 OR 114977-28-5/RN

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L3 183161 (HEPATOCELLULAR OR HEPATIC OR LIVER OR HEPATO) (W) (CANCER OR NEOPLASM OR NEOPLASTIC OR TUMOR OR TUMOUR OR CANCEROUS)

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PROCESSING COMPLETED FOR L4
L5 403 DUP REM L4 (31 DUPLICATES REMOVED).

=> focus
PROCESSING COMPLETED FOR L5
L6 403 FOCUS L5 1-

=> s (hepatocellular or hepatic or liver or hepat) (w) (cancer or neoplasm or neoplastic or tumor or tumour or cancerous or carcinoma or carci?)
L7 231795 (HEPATOCELLULAR OR HEPATIC OR LIVER OR HEPATO) (W) (CANCER OR NEOPLASM OR NEOPLASTIC OR TUMOR OR TUMOUR OR CANCEROUS OR CARCINOMA OR CARCI?)

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NEWS 6 SEP 11 CA/CAplus enhanced with more pre-1907 records
NEWS 7 SEP 21 CA/CAplus fields enhanced with simultaneous left and right truncation
NEWS 8 SEP 25 CA(SM)/CAplus(SM) display of CA Lexicon enhanced
NEWS 9 SEP 25 CAS REGISTRY(SM) no longer includes Concord 3D coordinates
NEWS 10 SEP 25 CAS REGISTRY(SM) updated with amino acid codes for pyrrolysine
NEWS 11 SEP 28 CEABA-VTB classification code fields reloaded with new classification scheme
NEWS 12 OCT 19 LOGOFF HOLD duration extended to 120 minutes
NEWS 13 OCT 19 E-mail format enhanced
NEWS 14 OCT 23 Option to turn off MARPAT highlighting enhancements available
NEWS 15 OCT 23 CAS Registry Number crossover limit increased to 300,000 in multiple databases
NEWS 16 OCT 23 The Derwent World Patents Index suite of databases on STN has been enhanced and reloaded
NEWS 17 OCT 30 CHEMLIST enhanced with new search and display field
NEWS 18 NOV 03 JAPIO enhanced with IPC 8 features and functionality
NEWS 19 NOV 10 CA/CAplus F-Term thesaurus enhanced
NEWS 20 NOV 10 STN Express with Discover! free maintenance release Version 8.01c now available
NEWS 21 NOV 13 CA/CAplus pre-1967 chemical substance index entries enhanced with preparation role
NEWS 22 NOV 20 CAS Registry Number crossover limit increased to 300,000 in additional databases
NEWS 23 NOV 20 CA/CAplus to MARPAT accession number crossover limit increased to 50,000
NEWS 24 NOV 20 CA/CAplus patent kind codes will be updated
NEWS 25 DEC 01 CAS REGISTRY updated with new ambiguity codes

NEWS EXPRESS NOVEMBER 10 CURRENT WINDOWS VERSION IS V8.01c, CURRENT MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP), AND CURRENT DISCOVER FILE IS DATED 25 SEPTEMBER 2006.

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L8          469 L2 AND L7

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L9          420 DUP REM L8 (49 DUPLICATES REMOVED)

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PROCESSING COMPLETED FOR L9
L10         420 FOCUS L9 1-

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  2 FILES SEARCHED...
L11         55 L10 AND PD<=2000

=> focus
PROCESSING COMPLETED FOR L11
L12         55 FOCUS L11 1-

=> d ibib abs hitstr 1-55

L12 ANSWER 1 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER:      2000:650912 CAPLUS
DOCUMENT NUMBER:       134:141449
TITLE:                 Comparison of 2-methoxyestradiol-induced,
                      docetaxel-induced, and paclitaxel-induced
                      apoptosis in hepatoma cells and its correlation with
                      reactive oxygen species
AUTHOR(S):             Lin, Heng-Liang; Liu, Tsung-Yun; Chau, Gar-Yang; Lui,
                      Wing-Yiu; Chi, Chin-Wen
CORPORATE SOURCE:      Institute of Pharmacology, National Yang-Ming
                      University, Taipei, Taiwan
SOURCE:                Cancer (New York) (2000), 89(5), 983-994
CODEN: CANCAR; ISSN: 0008-543X
PUBLISHER:             John Wiley & Sons, Inc.
DOCUMENT TYPE:          Journal
LANGUAGE:              English

AB Previously, the authors observed that paclitaxel treatment of hepatoma cells resulted in differential cytotoxicity. Whether other antimicrotubule agents (docetaxel and 2-methoxyestradiol) are more effective than paclitaxel is not clear. Moreover, whether the modulation of reactive oxygen species (ROS) is involved in the drug-induced growth inhibition of hepatoma cells is not known. The authors examined the effects of 2-methoxyestradiol, paclitaxel, and docetaxel on HepG2, Hep3B, HA22T/VGH, and Hepal-6 hepatoma cell lines. The parameters examined included cell viability, cell membrane permeability, cell cycle distribution, DNA fragmentation, and ROS generation. Docetaxel and paclitaxel inhibited the growth of hepatoma cells at submicromolar concns., whereas that of 2-methoxyestradiol was within a micromolar range. This drug-induced growth inhibition was cell cycle dependent. 2-Methoxyestradiol-treated (10-50 µM) cells resulted in G2/M block prior to apoptosis. High dose (0.1 µM) docetaxel- and paclitaxel-treated cells resulted in a G2/M arrest followed by generation of polyploidy or apoptosis; however, low dose (0.01 µM) treatment induced apoptosis without G2/M arrest. The low dose effect was more significant in docetaxel-treated cells than in paclitaxel-treated cells. Although these antimicrotubule agents increased the formation of ROS, antioxidant treatment did not block drug-induced cell cycle and growth inhibition effects. The current results suggest that the growth inhibition of hepatoma cells induced by 2-methoxyestradiol, paclitaxel, and docetaxel was mediated through G2/M-phase arrest, caspase activation, and DNA fragmentation. The drug-induced apoptosis was independent of ROS formation. Docetaxel was more effective than paclitaxel in killing hepatoma
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cells. The potential of using 2-methoxyestradiol and docetaxel for the treatment of patients with hepatoma is worthy of further study.

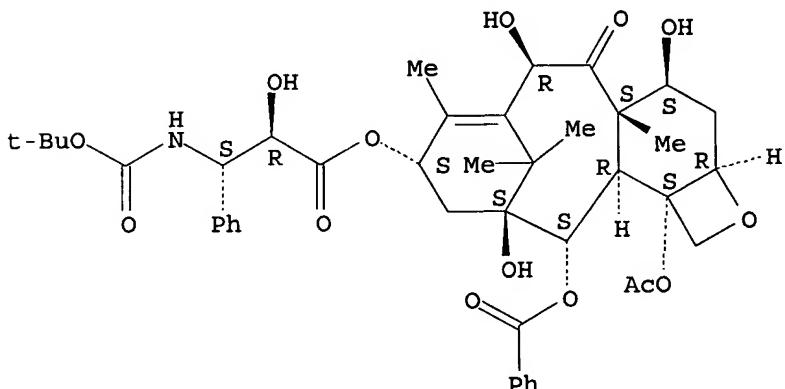
IT 114977-28-5, Docetaxel
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(2-methoxyestradiol-, docetaxel-, and paclitaxel-induced apoptosis in hepatoma cells)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 58 THERE ARE 58 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 2 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:126569 CAPLUS

DOCUMENT NUMBER: 132:175461

TITLE: Factors predicting for efficacy and safety of docetaxel in a compassionate-use cohort of 825 heavily pretreated advanced breast cancer patients

AUTHOR(S): Alexandre, J.; Bleuzen, P.; Bonneterre, J.; Sutherland, W.; Misset, J. L.; Guastalla, J.-P.; Viens, P.; Faivre, S.; Chahine, A.; Spielman, M.; Bensmaine, A.; Marty, M.; Mahjoubi, M.; Cvitkovic, E.

CORPORATE SOURCE: Paul Brousse Hospital and Institut Gustave Roussy, Villejuif, 94804, Fr.

SOURCE: Journal of Clinical Oncology (2000), 18(3), 562-573

CODEN: JCONDN; ISSN: 0732-183X

PUBLISHER: Lippincott Williams & Wilkins

DOCUMENT TYPE: Journal

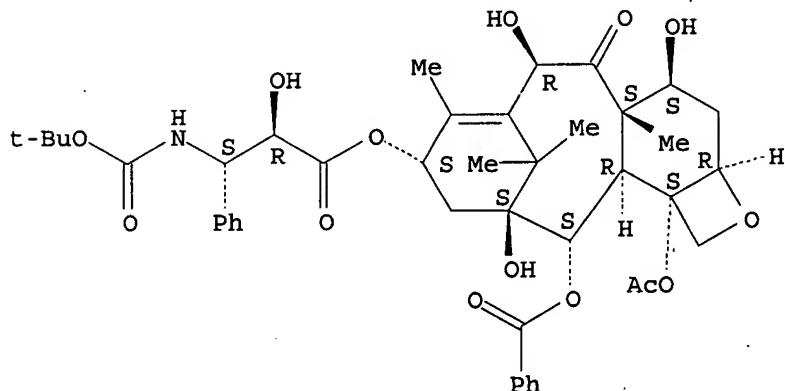
LANGUAGE: English

AB Purpose: To identify predictive factors for efficacy and safety in advanced breast cancer (ABC) patients treated in the French compassionate-use docetaxel program. Patients and Methods: A total of 825 ABC patients treated with docetaxel (100 mg/m² every 3 wk) were source-reviewed and analyzed for prognostic factors associated with overall response rate (ORR), time to treatment failure (TTF), overall survival (OS), febrile neutropenia, mucositis, and severe fluid retention syndrome by univariate and multivariate anal. Results: The ORR was 22.9% (95% confidence interval, 20.2% to 26.2%). The median TTF and

OS were 4.0 and 9.8 mo, resp. By multivariate anal., secondary anthracycline-resistant disease was significantly associated ($P < .05$) with lower ORR and shorter TTF and OS, whereas anthracycline-refractory disease was associated with shorter OS. Poor performance status was associated with lower ORR, shorter TTF, and shorter OS. Liver dysfunction (transaminase levels > 1.5 times the upper limit of normal [ULN] and alkaline phosphatase [AP] level $>$ three times ULN) and time since first relapse less than 24 mo were associated with shorter TTF and OS. Other significant correlations included the following: elevated CA 15-3 serum level with lower ORR; more than two involved sites, and minor transaminase and AP level abnormalities with shorter OS; and no previous chemotherapy for ABC with shorter TTF. According to multivariate anal., ORR, TTF, and OS were not decreased in patients with liver metastases but without liver dysfunction. Conclusion: Docetaxel activity was maintained in heavily pretreated ABC patients and in those with liver metastasis; docetaxel must be used cautiously, however, in patients with liver dysfunction in whom high morbidity risk necessitates strict adherence to dose-adaptation guidelines.

IT 114977-28-5, Docetaxel
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (factors predicting for efficacy and safety of docetaxel in
 pretreated humans with advanced breast cancer)
 RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]-
 α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-
 (benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-
 trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-
 cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA
 INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 40 THERE ARE 40 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 3 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2000:60933 CAPLUS
 DOCUMENT NUMBER: 132:102535
 TITLE: A Phase I study of gemcitabine and docetaxel
 in patients with metastatic solid tumors
 AUTHOR(S): Ryan, David P.; Lynch, Thomas J.; Grossbard, Michael
 L.; Seiden, Michael V.; Fuchs, Charles S.; Grenon,
 Nina; Baccala, Paul; Berg, Deborah; Finkelstein,
 Dianne; Mayer, Robert J.; Clark, Jeffrey W.
 CORPORATE SOURCE: Gastrointestinal Cancer Clinic, Dana-Farber/Partners
 CancerCare, Boston, MA, 02114, USA
 SOURCE: Cancer (New York) (2000), 88(1), 180-185
 CODEN: CANCAR; ISSN: 0008-543X

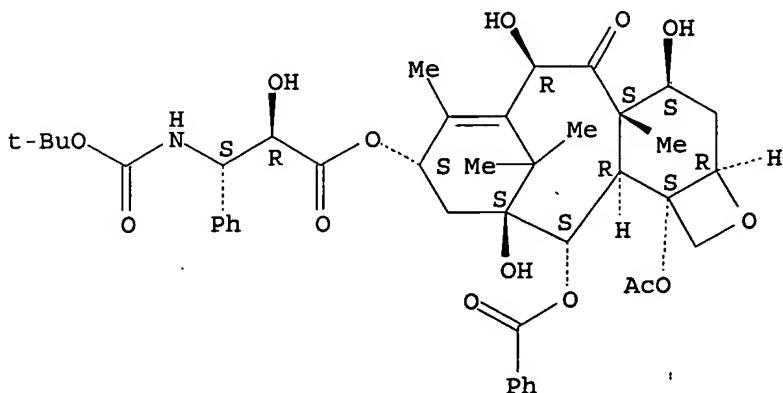
PUBLISHER: John Wiley & Sons, Inc.
 DOCUMENT TYPE: Journal
 LANGUAGE: English

AB A Phase I study was initiated to determine the maximum tolerated dose of weekly gemcitabine combined with monthly, fixed-dose docetaxel. Patients with metastatic solid tumors were treated with docetaxel, 60 mg/m², on Day 1 every 28 days. Gemcitabine was administered on Days 1, 8, and 15 and underwent dose adjustment in cohorts of 3-6 patients. At the maximum tolerated dose, 11 addnl. patients were enrolled. Twenty-six patients received 85 cycles of therapy. At the first dose level, the planned gemcitabine dose on Days 1, 8, and 15 was 800 mg/m². Two of the 6 patients treated at this dose level experienced dose-limiting toxicities (DLTs) requiring the reduction of gemcitabine to 600 mg/m² per dose and the administration of ciprofloxacin, 500 mg orally twice daily, on Days 8-18. At the second dose level the first 3 patients experienced no DLTs and the dose of gemcitabine was increased to 700 mg/m². Two of the 6 patients treated at the 700 mg/m² dose level experienced DLTs. Eleven addnl. patients were enrolled at the recommended Phase II dose of gemcitabine (600 mg/m²). At this dose level, Grade 3/4 (according to the National Cancer Institute's common toxicity criteria) neutropenia and thrombocytopenia occurred in 12.5% and 2.1% of cycles, resp. Grade 3 and 4 nonhematol. toxicities were uncommon. Three of seven evaluable patients with pancreatic carcinoma had evidence of significant antineoplastic activity (three partial responses). In addition, two complete responses (one patient with gastric carcinoma and one patient with ovarian carcinoma) and one partial response (patient with hepatocellular carcinoma) were noted in patients with other solid tumors. The regimen comprised of docetaxel, 60 mg/m², on Day 1 and gemcitabine, 600 mg/m², on Days 1, 8, and 15 with ciprofloxacin on Days 8-18 every 28 days is safe, well tolerated, and active.

IT 114977-28-5, Docetaxel
 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (gemcitabine and docetaxel in human patients with metastatic solid tumors)

RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



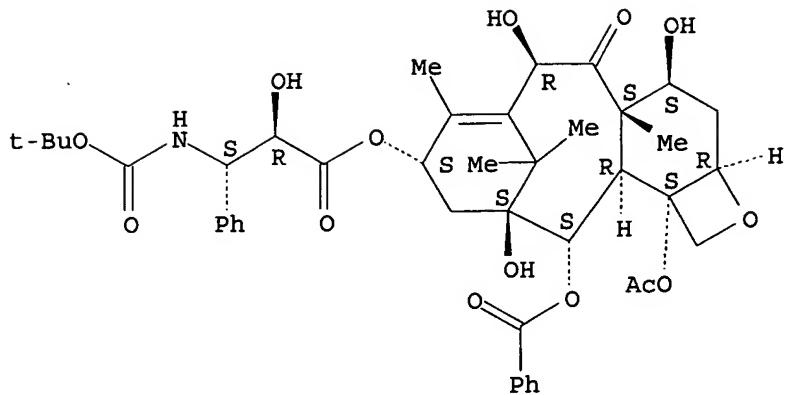
L12 ANSWER 4 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 1999:641123 CAPLUS
DOCUMENT NUMBER: 132:117128
TITLE: Efficacy and safety of docetaxel (Taxotere)
in heavily pretreated advanced breast cancer patients;
the French compassionate use program experience
AUTHOR(S): Bonneterre, J.; Spielman, M.; Guastalla, J. -P.;
Marty, M.; Viens, P.; Chollet, P.; Roche, H.;
Fumoleau, P.; Mauriac, L.; Bourgeois, H.; Namer, M.;
Bergerat, J. P.; Misset, J. -L.; Trandafir, L.;
Mahjoubi, M.
CORPORATE SOURCE: Centre Oscar Lambret, Lille, 59020, Fr.
SOURCE: European Journal of Cancer (1999), 35(10),
1431-1439
CODEN: EJCAEL; ISSN: 0959-8049
PUBLISHER: Elsevier Science Ltd.
DOCUMENT TYPE: Journal
LANGUAGE: English

AB The aim was to assess retrospectively doceaxel safety and efficacy in advanced breast cancer patients in a French compassionate use program. Patients had received >1 prior chemotherapy regimen for advanced disease, were either anthracycline-resistant (that is progressed within 6 mo after anthracycline-based chemotherapy) or had received the maximum cumulative dose. The recommended docetaxel dose was 100 mg/m²/cycle (75 mg/m²) prior palliative chemotherapy lines. The most frequent severe toxicity, febrile neutropenia (reported in 223/870 (25.6%) patients evaluable for safety), caused 10 deaths, 6 of these being patients with severe liver impairment before inclusion. Fluid retention syndrome and other common non-Hematol. toxicities were well tolerated. 3.1% (28/889) of all patients and 11.4% of those with liver dysfunction, died from treatment-related causes. The overall response rate in 825 assessable patients was 22.9% (95% confidence interval (CI): 20.2-26.2%). Median time to treatment failure was 4 mo (95% CI: 3.6-4.3) and median survival was 9.8 mo (95% CI: 8.8-10.7). This report on the largest series of unselected advanced breast cancer patients treated with docetaxel , supports previous phase II studies, confirming docetaxel's utility in patients relapsing after failing anthracycline-containing palliative chemotherapy.

IT 114977-28-5, Docetaxel
RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(efficacy and safety of docetaxel in heavily pretreated advanced breast cancer patients)

RN 114977-28-5 CAPLUS
CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 5 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2000:790283 CAPLUS
 DOCUMENT NUMBER: 133:344606
 TITLE: Combined pharmaceuticals comprising anthracycline derivatives
 INVENTOR(S): Geroni, Maria Cristina; Ripamonti, Marina; Caruso, Michele; Suarato, Antonino
 PATENT ASSIGNEE(S): Pharmacia & Upjohn S.p.A., Italy
 SOURCE: PCT Int. Appl., 23 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|------------------|--------------|
| WO 2000066093 | A2 | 20001109 | WO 2000-EP2923 | 20000404 <-- |
| WO 2000066093 | A3 | 20010125 | | |
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| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| EP 1173187 | A2 | 20020123 | EP 2000-925158 | 20000404 |
| EP 1173187 | B1 | 20030806 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| JP 2002543112 | T2 | 20021217 | JP 2000-614978 | 20000404 |
| EP 1323423 | A1 | 20030702 | EP 2003-75776 | 20000404 |
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| AT 246507 | E | 20030815 | AT 2000-925158 | 20000404 |
| PT 1173187 | T | 20031031 | PT 2000-925158 | 20000404 |
| ES 2204572 | T3 | 20040501 | ES 2000-925158 | 20000404 |
| CN 1507869 | A | 20040630 | CN 2003-10114911 | 20000404 |
| CN 1535688 | A | 20041013 | CN 2003-10114909 | 20000404 |
| CN 1853645 | A | 20061101 | CN 2006-10059701 | 20000404 |
| CN 1853642 | A | 20061101 | CN 2006-10059703 | 20000404 |
| TW 222863 | B1 | 20041101 | TW 2000-89106805 | 20000412 |
| US 6537990 | B1 | 20030325 | US 2001-926392 | 20011025 |
| HK 1045462 | A1 | 20060428 | HK 2002-107029 | 20020926 |

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|------------------------|-------------|------------------|-------------|
| US 2003087839 | A1 20030508 | US 2002-284144 | 20021031 |
| US 6586428 | B2 20030701 | | |
| PRIORITY APPLN. INFO.: | | GB 1999-9925 | A 19990429 |
| | | CN 2000-806897 | A 20000404 |
| | | CN 2003-10114909 | A3 20000404 |
| | | CN 2003-10114911 | A3 20000404 |
| | | EP 2000-925158 | A3 20000404 |
| | | WO 2000-EP2923 | W 20000404 |
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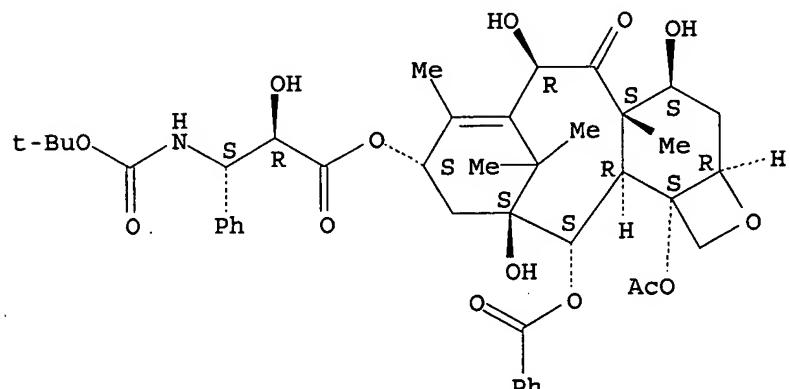
AB The present invention relates to combined pharmaceuticals comprising a morpholinylanthracycline administered in combination anticancer agents chosen from an alkylating agent, an antimetabolite, a topoisomerase II inhibitor, a topoisomerase I inhibitor, an antimitotic drug and a platinum derivative, which are useful in anticancer therapy, particularly in the treatment of a primary or metastatic liver cancer. At doses 5.9 and 7.7 mg/kg cis-platin administered alone and 0.05 mg/kg PNU-152243 (morpholinylanthracycline) administered alone, the increase in lifespan values was 33, 33 and 50%. Sequential administration of these drugs showed a therapeutic advantage of the combination in comparison with each drug administered alone.

IT 114977-28-5, Docetaxel
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (combined pharmaceuticals comprising anthracycline derivs.)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 6 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:456927 CAPLUS

DOCUMENT NUMBER: 133:84243

TITLE: Method of using a cyclooxygenase-2 inhibitor and one or more antineoplastic agents as a combination therapy in the treatment of neoplasia

INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.

PATENT ASSIGNEE(S): G.D. Searle and Co., USA

SOURCE: PCT Int. Appl., 236 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 21

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-------------------|--------------|
| WO 2000038730 | A2 | 20000706 | WO 1999-US30693 | 19991222 <-- |
| WO 2000038730 | A3 | 20001102 | | |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2356606 | AA | 20000706 | CA 1999-2356606 | 19991222 <-- |
| AU 2000023805 | A5 | 20000731 | AU 2000-23805 | 19991222 <-- |
| AU 783992 | B2 | 20060112 | | |
| EP 1140192 | A2 | 20011010 | EP 1999-967543 | 19991222 |
| EP 1140192 | B1 | 20060405 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, CY | | | | |
| TR 200102499 | T2 | 20011221 | TR 2001-200102499 | 19991222 |
| BR 9916518 | A | 20020129 | BR 1999-16518 | 19991222 |
| HU 200104814 | A2 | 20020429 | HU 2001-4814 | 19991222 |
| JP 2002533416 | T2 | 20021008 | JP 2000-590681 | 19991222 |
| EP 1522313 | A1 | 20050413 | EP 2004-26577 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, RO, CY | | | | |
| AT 322290 | E | 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A | 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A | 20020107 | ZA 2001-5120 | 20010621 |
| NO 2001003155 | A | 20010822 | NO 2001-3155 | 20010622 |
| US 2003119895 | A1 | 20030626 | US 2002-150546 | 20020516 |
| US 2003203956 | A1 | 20031030 | US 2002-212523 | 20020805 |
| AU 2004210578 | A1 | 20041007 | AU 2004-210578 | 20040910 |
| US 2005037090 | A1 | 20050217 | US 2004-945422 | 20040920 |
| PRIORITY APPLN. INFO.: | | | US 1998-113786P | P 19981223 |
| | | | US 1999-385214 | A 19990827 |
| | | | AU 2000-25936 | A3 19991222 |
| | | | EP 1999-968939 | A3 19991222 |
| | | | WO 1999-US30693 | W 19991222 |
| | | | US 2001-857873 | A2 20011005 |

AB Methods are provided to treat or prevent neoplasia disorders in a mammal using a combination of a cyclooxygenase-2 inhibitor and an antineoplastic agent.

IT 114977-28-5, Docetaxel

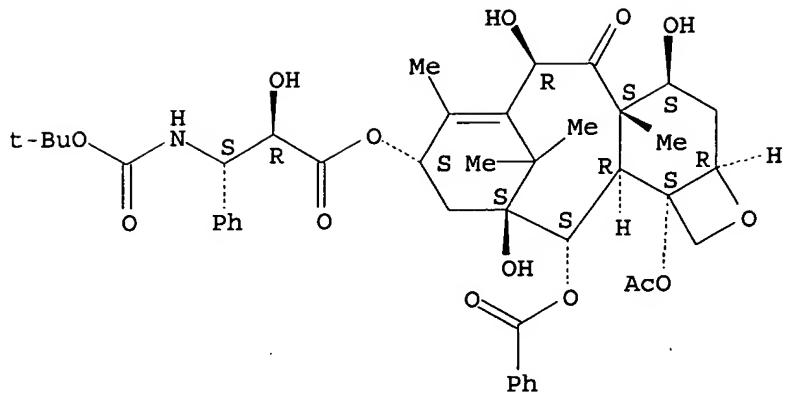
RL: BAC (Biological activity or effector; except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(cyclooxygenase-2 inhibitor-antineoplastic agent combination for neoplasia treatment)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[1,1-dimethylethoxy carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

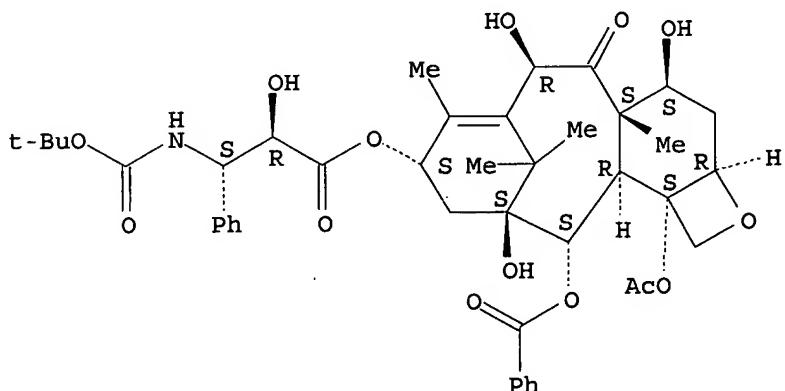


L12 ANSWER 7 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2000:456915 CAPLUS
 DOCUMENT NUMBER: 133:84242
 TITLE: Method of using a matrix-metalloproteinase inhibitor and one or more antineoplastic agents as a combination therapy in the treatment of neoplasia
 INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.
 PATENT ASSIGNEE(S): G.D. Searle and Co., USA
 SOURCE: PCT Int. Appl., 277 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 21
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-------------------|--------------|
| WO 2000038718 | A2 | 20000706 | WO 1999-US30699 | 19991222 <-- |
| WO 2000038718 | A3 | 20001109 | | |
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| AU 2000027135 | A5 | 20000731 | AU 2000-27135 | 19991222 <-- |
| EP 1140182 | A2 | 20011010 | EP 1999-968941 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| TR 200102499 | T2 | 20011221 | TR 2001-200102499 | 19991222 |
| JP 2002533406 | T2 | 20021008 | JP 2000-590669 | 19991222 |
| EP 1522313 | A1 | 20050413 | EP 2004-26577 | 19991222 |
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| AT 322290 | E | 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A | 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A | 20020107 | ZA 2001-5120 | 20010621 |
| US 6858598 | B1 | 20050222 | US 2001-857995 | 20011005 |
| AU 2004210578 | A1 | 20041007 | AU 2004-210578 | 20040910 |
| US 2005058725 | A1 | 20050317 | US 2004-945002 | 20040920 |
| US 6916800 | B2 | 20050712 | | |
| PRIORITY APPLN. INFO.: | | | US 1998-113786P | P 19981223 |

| | | | |
|----|---|-----------------|-------------|
| | | US 1999-385214 | A 19990827 |
| | | AU 2000-25936 | A3 19991222 |
| | | EP 1999-968939 | A3 19991222 |
| | | WO 1999-US30699 | W 19991222 |
| | | US 2001-857995 | A1 20011005 |
| AB | Methods are provided for the prevention and treatment of neoplasia disorders in a mammal using a combination of a matrix metalloproteinase inhibitor and an antineoplastic agent. | | |
| IT | 114977-28-5, Docetaxel
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(matrix metalloproteinase inhibitor and antineoplastic agent as combination therapy in neoplasia treatment) | | |
| RN | 114977-28-5 CAPLUS | | |
| CN | Benzene propanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME) | | |

Absolute stereochemistry.



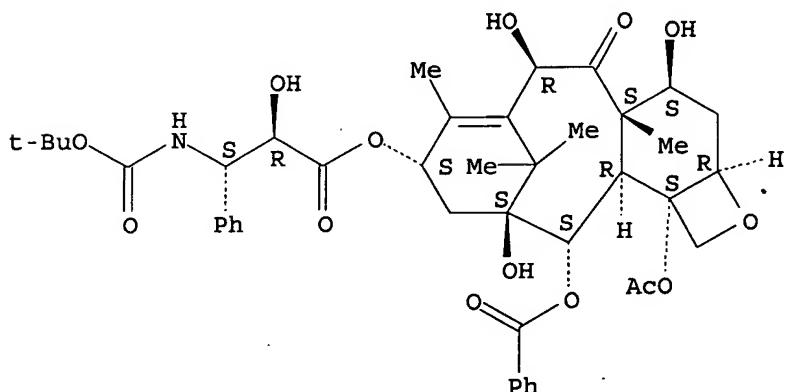
L12 ANSWER 8 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2000:456866 CAPLUS
 DOCUMENT NUMBER: 133:84239
 TITLE: Method of using an integrin antagonist and one or more antineoplastic agents as a combination therapy in the treatment of neoplasia
 INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.
 PATENT ASSIGNEE(S): G.D. Searle and Co., USA
 SOURCE: PCT Int. Appl., 220 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 21
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|-----------------|-------------|
| WO 2000038665 | A2 | 20000706 | WO 1999-US30670 | 19991222 <- |
| WO 2000038665 | A3 | 20001116 | | |
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 MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
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 TR 200102499 T2 20011221 TR 2001-200102499 19991222
 JP 2002533387 T2 20021008 JP 2000-590619 19991222
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 AT 322290 E 20060415 AT 1999-967543 19991222
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 ZA 2001005120 A 20020107 ZA 2001-5120 20010621
 US 6833373 B1 20041221 US 2001-857994 20011005
 US 2004234624 A1 20041125 US 2004-865414 20040610
 AU 2004210578 A1 20041007 AU 2004-210578 20040910
 PRIORITY APPLN. INFO.: US 1998-113786P P 19981223
 US 1999-385214 A 19990827
 AU 2000-25936 A3 19991222
 EP 1999-968939 A3 19991222
 WO 1999-US30670 W 19991222
 US 2001-857994 A1 20011005

- AB The present invention provides methods to treat or prevent neoplasia disorders in a mammal using a combination of an integrin antagonist and an antineoplastic agent.
 IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (integrin antagonist-antineoplastic agent combination for neoplasia treatment)
 RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -{[(1,1-dimethylethoxy)carbonyl]amino}- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S) - (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 9 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2000:368032 CAPLUS
 DOCUMENT NUMBER: 133:26843
 TITLE: Methods and compositions for diagnosis and treatment
 of cancer based on the transcription factor ets2
 INVENTOR(S): Papas, Takis S.; Watson, Dennis K.
 PATENT ASSIGNEE(S): Musc Foundation for Research Development, USA; Papas,
 Tula Christy
 SOURCE: PCT Int. Appl., 101 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 2000030590 | A2 | 20000602 | WO 1999-US27805 | 19991123 <-- |
| WO 2000030590 | A3 | 20000817 | | |
| W: AU, CA, JP, US | | | | |
| RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
PT, SE | | | | |
| CA 2351627 | AA | 20000602 | CA 1999-2351627 | 19991123 <-- |
| AU 2000024740 | A5 | 20000613 | AU 2000-24740 | 19991123 <-- |
| EP 1133575 | A2 | 20010919 | EP 1999-968046 | 19991123 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI | | | | |
| JP 2002530102 | T2 | 20020917 | JP 2000-583475 | 19991123 |
| US 2002081601 | A1 | 20020627 | US 2001-841963 | 20010425 |
| US 2004047845 | A1 | 20040311 | US 2001-841960 | 20010425 |
| PRIORITY APPLN. INFO.: | | | US 1998-109850P | P 19981125 |
| | | | WO 1999-US27805 | W 19991123 |

AB The present invention relates to methods for treating and preventing
 cancer by modifying the expression of ets2 gene expression or the activity
 of the gene product. The invention also relates to sensitizing cancer
 cells to chemotherapeutic or radiotherapeutic agents. Ets2 gene
 expression and/or activity of the gene product can be modulated using
 antisense ets2 nucleic acids and/or modified ets2 proteins. The present
 invention also provides pharmaceutical compns. which comprise antisense
 ets2 nucleic acid, and nucleic acid that encode modified ets2 proteins
 and/or modified ets2 proteins.

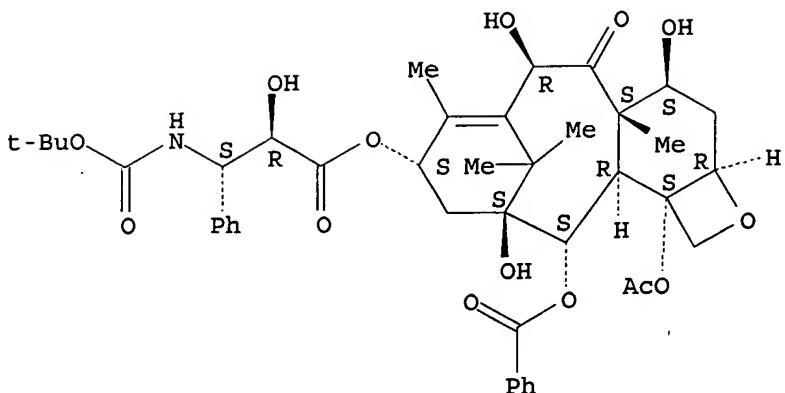
IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological
 study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
 (Uses)

(diagnosis and treatment of cancer based on the transcription factor
 ets2)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]-
 α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-
(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-
trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-
cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA
INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 10 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:456950 CAPLUS

DOCUMENT NUMBER: 133:84244

TITLE: Method of using a cyclooxygenase-2 inhibitor and an integrin antagonist as a combination therapy in the treatment of neoplasia

INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.

PATENT ASSIGNEE(S): G.D. Searle and Co., USA

SOURCE: PCT Int. Appl., 348 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

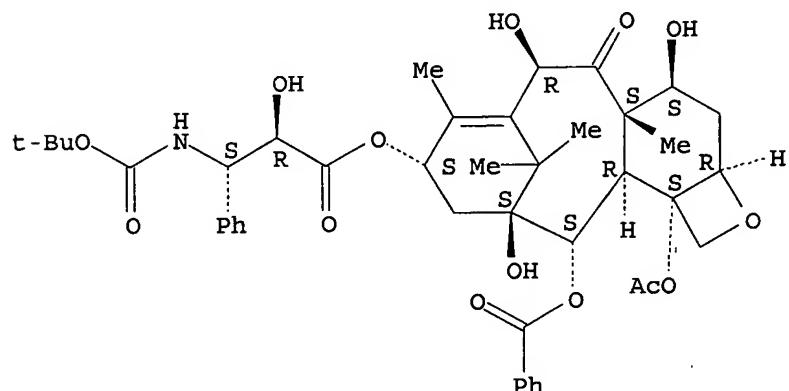
FAMILY ACC. NUM. COUNT: 21

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-------------------|--------------|
| WO 2000038786 | A2 | 20000706 | WO 1999-US30692 | 19991222 <-- |
| WO 2000038786 | A3 | 20010308 | | |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2356302 | AA | 20000706 | CA 1999-2356302 | 19991222 <-- |
| AU 2000022104 | A5 | 20000731 | AU 2000-22104 | 19991222 <-- |
| EP 1140179 | A2 | 20011010 | EP 1999-966594 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| TR 200102499 | T2 | 20011221 | TR 2001-200102499 | 19991222 |
| JP 2002533422 | T2 | 20021008 | JP 2000-590734 | 19991222 |
| EP 1522313 | A1 | 20050413 | EP 2004-26577 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, RO, CY | | | | |
| AT 322290 | E | 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A | 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A | 20020107 | ZA 2001-5120 | 20010621 |
| AU 2004210578 | A1 | 20041007 | AU 2004-210578 | 20040910 |
| PRIORITY APPLN. INFO.: | | | US 1998-113786P | P 19981223 |
| | | | US 1999-385214 | A 19990827 |
| | | | AU 2000-25936 | A3 19991222 |
| | | | EP 1999-968939 | A3 19991222 |

| | |
|----|---|
| AB | Methods are provided to treat or prevent neoplasia disorders in a mammal using a combination of a cyclooxygenase-2 inhibitor, an integrin antagonist and an antineoplastic agent. |
| IT | 114977-28-5, Docetaxel
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(cyclooxygenase-2 inhibitor and integrin antagonist in combination for neoplasia treatment) |
| RN | 114977-28-5 CAPLUS |
| CN | Benzene propanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME) |

Absolute stereochemistry.



L12 ANSWER 11 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2000:456916 CAPLUS
DOCUMENT NUMBER: 133:68929
TITLE: Use of a matrix metalloproteinase inhibitor and an integrin antagonist in the treatment of neoplasia
INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.; Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.
PATENT ASSIGNEE(S): G.D. Searle and Co., USA
SOURCE: PCT Int. Appl., 358 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 21
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---------------|--|----------|-----------------|--------------|
| WO 2000038719 | A1 | 20000706 | WO 1999-US30700 | 19991222 <-- |
| W: | AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL,
IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA,
MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI,
SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | |
| RW: | GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |

| | | | |
|--|-------------|-------------------|--------------|
| CA 2356402 | AA 20000706 | CA 1999-2356402 | 19991222 <-- |
| AU 2000027136 | A5 20000731 | AU 2000-27136 | 19991222 <-- |
| EP 1140183 | A1 20011010 | EP 1999-968942 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO | | | |
| TR 200102499 | T2 20011221 | TR 2001-200102499 | 19991222 |
| JP 2002533407 | T2 20021008 | JP 2000-590670 | 19991222 |
| EP 1522313 | A1 20050413 | EP 2004-26577 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI, RO, CY | | | |
| AT 322290 | E 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A 20020107 | ZA 2001-5120 | 20010621 |
| AU 2004210578 | A1 20041007 | AU 2004-210578 | 20040910 |
| PRIORITY APPLN. INFO.: | | | |
| US 1998-113786P P 19981223 | | | |
| US 1999-385214 A 19990827 | | | |
| AU 2000-25936 A3 19991222 | | | |
| EP 1999-968939 A3 19991222 | | | |
| WO 1999-US30700 W 19991222 | | | |

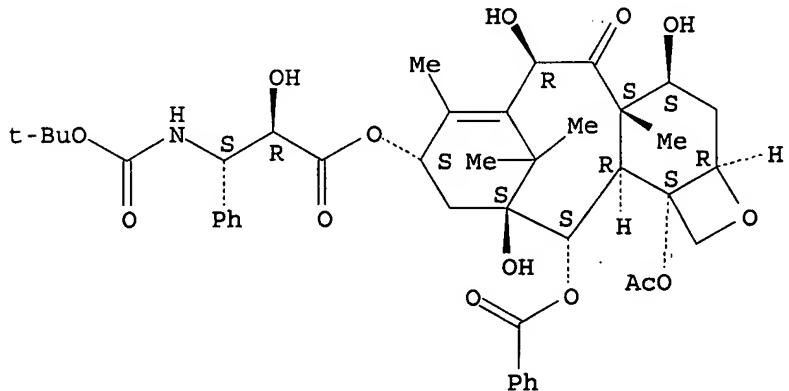
AB Methods are provided to treat or prevent neoplasia disorders in a mammal using a combination of a matrix metalloproteinase inhibitor, an integrin antagonist, and an antineoplastic agent.

IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (matrix metalloproteinase inhibitor and integrin antagonist in neoplasia treatment)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 12 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:441655 CAPLUS

DOCUMENT NUMBER: 133:68922

TITLE: Method of using a cyclooxygenase-2 inhibitor and a matrix metalloproteinase inhibitor as a combination therapy in the treatment of neoplasia

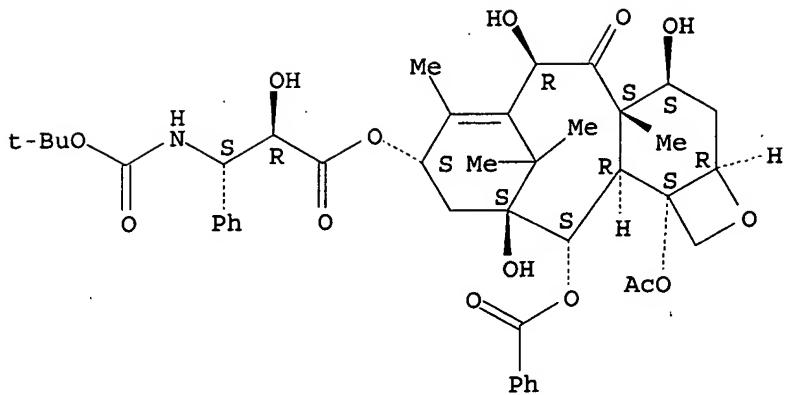
INVENTOR(S): McKearn, John P.; Gordon, Gary; Cunningham, James J.;

Gately, Stephen T.; Koki, Alane T.; Masferrer, Jaime L.
 PATENT ASSIGNEE(S): G.D. Searle and Co., USA
 SOURCE: PCT Int. Appl., 437 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 21
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-------------------|--------------|
| WO 2000037107 | A2 | 20000629 | WO 1999-US30776 | 19991222 <-- |
| WO 2000037107 | A3 | 20010201 | | |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2356426 | AA | 20000629 | CA 1999-2356426 | 19991222 <-- |
| AU 2000025936 | A5 | 20000712 | AU 2000-25936 | 19991222 <-- |
| EP 1140194 | A2 | 20011010 | EP 1999-968540 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| TR 200102499 | T2 | 20011221 | TR 2001-200102499 | 19991222 |
| BR 9916536 | A | 20020102 | BR 1999-16536 | 19991222 |
| HU 200104747 | A2 | 20020429 | HU 2001-4747 | 19991222 |
| JP 2002532563 | T2 | 20021002 | JP 2000-589217 | 19991222 |
| EP 1522313 | A1 | 20050413 | EP 2004-26577 | 19991222 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI, RO, CY | | | | |
| AT 322290 | E | 20060415 | AT 1999-967543 | 19991222 |
| ZA 2001005055 | A | 20020920 | ZA 2001-5055 | 20010620 |
| ZA 2001005120 | A | 20020107 | ZA 2001-5120 | 20010621 |
| NO 2001003156 | A | 20010823 | NO 2001-3156 | 20010622 |
| AU 2004210578 | A1 | 20041007 | AU 2004-210578 | 20040910 |
| PRIORITY APPLN. INFO.: | | | US 1998-113786P | P 19981223 |
| | | | US 1999-385214 | A 19990827 |
| | | | AU 2000-25936 | A3 19991222 |
| | | | EP 1999-968939 | A3 19991222 |
| | | | WO 1999-US30776 | W 19991222 |

- AB Methods are provided to treat or prevent neoplasia disorders in a mammal using a combination of a cyclooxygenase-2 inhibitor, a matrix metalloproteinase inhibitor and an antineoplastic agent.
 IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (cyclooxygenase-2 inhibitor and matrix metalloproteinase inhibitor in combination therapy for neoplasia treatment)
 RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -[[{(1,1-dimethylethoxy)carbonyl]amino}- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 13 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1997:400101 CAPLUS

DOCUMENT NUMBER: 127:23742

TITLE: Method, compositions and kits for increasing the oral bioavailability of pharmaceutical agents

INVENTOR(S): Broder, Samuel; Duchin, Kenneth L.; Selim, Sami

PATENT ASSIGNEE(S): Baker Norton Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 136 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 4

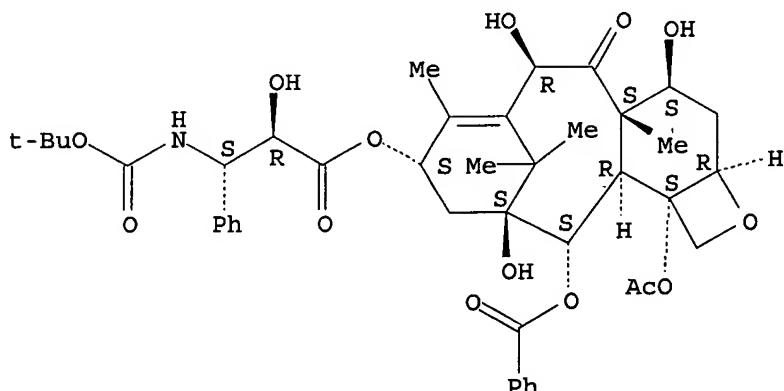
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 9715269 | A2 | 19970501 | WO 1996-IB1485 | 19961024 <-- |
| WO 9715269 | A3 | 19970731 | | |
| W: AM, AT, AU, BB, BG, BR, BY, CA, CH, CN, CZ, DE, DK, EE, ES, FI, GB, GE, HU, IL, JP, KE, KG, KP, KR, KZ, LK, LR, LT, LU, LV, MD, MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, TJ, TT, UA, UZ, VN | | | | |
| RW: KE, LS, MW, SD, SZ, UG, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
| US 5968972 | A | 19991019 | US 1996-608776 | 19960229 <-- |
| US 6245805 | B1 | 20010612 | US 1996-733142 | 19961016 |
| AU 9712056 | A1 | 19970515 | AU 1997-12056 | 19961024 <-- |
| AU 698142 | B2 | 19981022 | | |
| EP 794794 | A1 | 19970917 | EP 1996-943268 | 19961024 <-- |
| EP 794794 | B1 | 20051207 | | |
| R: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE | | | | |
| JP 10509741 | T2 | 19980922 | JP 1997-516449 | 19961024 <-- |
| JP 3361102 | B2 | 20030107 | | |
| BR 9607066 | A | 20021210 | BR 1996-7066 | 19961024 |
| RU 2217135 | C2 | 20031127 | RU 1997-112888 | 19961024 |
| PL 188281 | B1 | 20050131 | PL 1996-321791 | 19961024 |
| AT 311903 | E | 20051215 | AT 1996-943268 | 19961024 |
| ZA 9609001 | A | 19970617 | ZA 1996-9001 | 19961025 <-- |
| NO 9702968 | A | 19970723 | NO 1997-2968 | 19970625 <-- |
| NO 321091 | B1 | 20060313 | | |
| HK 1001960 | A1 | 20060127 | HK 1998-101042 | 19980211 |
| AU 2002035584 | A5 | 20020606 | AU 2002-35584 | 20020422 |
| AU 784159 | B2 | 20060216 | | |
| PRIORITY APPLN. INFO.: | | | US 1995-7071P | P 19951026 |
| | | | US 1996-608776 | A 19960229 |

US 1996-733142 A 19961016
 WO 1996-IB1485 W 19961024
 AU 1998-71300 A3 19980422

- AB A method of increasing the bioavailability upon oral administration of a pharmacol. active target agent, particularly an antitumor or antineoplastic agent which exhibits poor or inconsistent oral bioavailability (e.g., paclitaxel, docetaxel or etoposide), comprises the oral co-administration to a mammalian patient of the target agent and an oral bioavailability-enhancing agent (e.g., cyclosporin A, cyclosporin D, cyclosporin F, or ketoconazole). The oral bioavailability-enhancing agents are known to be MDR (P-glycoprotein) inhibitors. The enhancing agent may be administered orally from 0.5-24 h prior to the oral administration of one or more doses of the target agent, substantially simultaneously with the target agent, or both prior to and substantially simultaneously with the target agent. A method of treating mammalian patients suffering from diseases responsive to target agents with poor oral bioavailability, as well as oral dosage forms containing such target agents, combination oral dosage forms containing bioavailability-enhancing agents and target agents kits containing enhancing and target agent dosage forms and dosing information for the co-administration of the same are also disclosed.
- IT 114977-28-5, Docetaxel
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (target; increasing oral bioavailability of pharmaceutical agents)
- RN 114977-28-5 CAPLUS
- CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 14 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

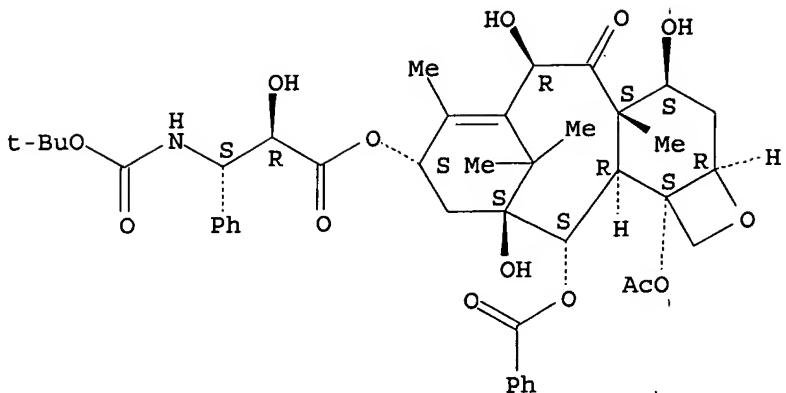
ACCESSION NUMBER: 1998:799989 CAPLUS
 DOCUMENT NUMBER: 130:43304
 TITLE: Method and compositions for administering taxanes orally to human patients using a cyclosporin to enhance bioavailability
 INVENTOR(S): Broder, Samuel; Duchin, Kenneth L.; Selim, Sami
 PATENT ASSIGNEE(S): Baker Norton Pharmaceuticals, Inc., USA
 SOURCE: PCT Int. Appl., 44 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 4

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------------|------------------|--------------|
| WO 9853811 | A1 | 19981203 | WO 1998-US7776 | 19980422 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG,
KP, KR, KZ, LC, LK, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX,
NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT,
UA, UG, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
| CA 2290446 | AA | 19981203 | CA 1998-2290446 | 19980422 <-- |
| AU 9871300 | A1 | 19981230 | AU 1998-71300 | 19980422 <-- |
| EP 994706 | A1 | 20000426 | EP 1998-918361 | 19980422 <-- |
| EP 994706 | B1 | 20051102 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI, RO | | | | |
| BR 9809694 | A | 20001003 | BR 1998-9694 | 19980422 <-- |
| JP 2002500667 | T2 | 20020108 | JP 1999-500663 | 19980422 |
| HU 200003546 | A2 | 20021128 | HU 2000-3546 | 19980422 |
| RU 2205005 | C2 | 20030527 | RU 1999-128033 | 19980422 |
| NZ 516026 | A | 20030630 | NZ 1998-516026 | 19980422 |
| CN 1550231 | A | 20041201 | CN 2004-10030478 | 19980422 |
| AT 308365 | E | 20051115 | AT 1998-918361 | 19980422 |
| ES 2247690 | T3 | 20060301 | ES 1998-918361 | 19980422 |
| ZA 9804268 | A | 19990623 | ZA 1998-4268 | 19980520 <-- |
| HK 1026637 | A1 | 20060106 | HK 2000-105943 | 20000920 |
| AU 2002035584 | A5 | 20020606 | AU 2002-35584 | 20020422 |
| AU 784159 | B2 | 20060216 | | |
| PRIORITY APPLN. INFO.: | | | | |
| | | US 1997-863513 | A 19970527 | |
| | | AU 1998-71300 | A3 19980422 | |
| | | NZ 1998-501127 | A1 19980422 | |
| | | WO 1998-US7776 | W 19980422 | |

- AB Taxane antineoplastic agents which have heretofore exhibited poor or non-existent oral bioavailability are administered orally to human patients suffering from taxane-responsive disease conditions and made sufficiently bioavailable to achieve therapeutic blood levels. In a preferred embodiment, the taxane, preferably paclitaxel, is co-administered to the patient with an oral cyclosporin enhancing agent, preferably cyclosporin A. By one preferred method, a dose of oral enhancer is administered about 0.5-72 h before the taxane and a second dose of the enhancer and administered immediately before, together with or immediately after the taxane. A method of treating human patients suffering from taxane-responsive disease conditions is also provided, as well as a method for providing such treatment while preventing or reducing hypersensitivity and allergic reactions without the need for pre-medication.
- IT 114977-28-5, Docetaxel
 RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (method and compns. for administering taxanes orally to human patients using a cyclosporin to enhance bioavailability)
- RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -{[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)}

Absolute stereochemistry.



REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 15 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1999:215571 CAPLUS

DOCUMENT NUMBER: 130:247032

TITLE: Fused imidazole derivatives for improving oral bioavailability of pharmaceutical agents

INVENTOR(S): Snoeck, Henricus Johannes Matheus

PATENT ASSIGNEE(S): Janssen Pharmaceutica N.V., Belg.

SOURCE: PCT Int. Appl., 45 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

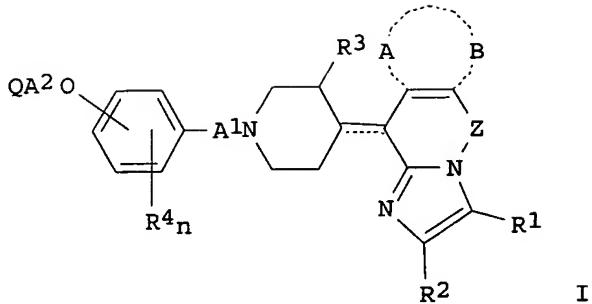
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 9913871 | A2 | 19990325 | WO 1998-EP5751 | 19980910 <-- |
| WO 9913871 | A3 | 19990603 | | |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 9911460 | A1 | 19990405 | AU 1999-11460 | 19980910 <-- |
| EP 1011726 | A2 | 20000628 | EP 1998-954268 | 19980910 <-- |
| EP 1011726 | B1 | 20030226 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, PT, IE, SI, LT, LV, FI, RO | | | | |
| JP 2001516716 | T2 | 20011002 | JP 2000-511494 | 19980910 |
| JP 3775780 | B2 | 20060517 | | |
| AT 233104 | E | 20030315 | AT 1998-954268 | 19980910 |
| ZA 9808527 | A | 20000322 | ZA 1998-8527 | 19980917 <-- |
| US 6544979 | B1 | 20030408 | US 2000-508748 | 20000315 |
| PRIORITY APPLN. INFO.: | | | EP 1997-202862 | A 19970918 |
| | | | WO 1998-EP5751 | W 19980910 |

OTHER SOURCE(S): MARPAT 130:247032

GI



AB Compds. I [dotted line = optional bond; n = 1, 2; R1 = H, halo, formyl, (substituted) C1-4 alkyl, etc.; R2 = H, halo, C1-4 alkyl, hydroxy-C1-4 alkyl, etc.; R3 = H, C1-4 alkyl, C1-4 alkyloxy; R4 = H, halo, C1-4 alkyl, C1-4 alkyloxy, halo-C1-4 alkyl; Z = CH₂, CH₂CH₂, CH=CH, CH(OH)CH₂, OCH₂, C(O)CH₂, C(=NOH)CH₂; AB = bivalent radical; Al = direct bond, (substituted) C1-6 alkanediyl, C1-6 alkanediyl-oxy-C1-6 alkanediyl, carbonyl, C1-6 alkanediylcarbonyl, (substituted) C1-6 alkanediylloxy; A2 = direct bond, C1-6 alkanediyl; Q = aryl], and N-oxide forms, pharmaceutically acceptable addition salts, and stereochem. isomeric forms thereof, are used for the manufacture of a medicine for improving the bioavailability of a second pharmaceutical agent which is co-administered orally to a warm-blooded animal. The second pharmaceutical agent is e.g. an antitumor agent. Preparation of compds. of the invention, and intermediates thereto, is described.

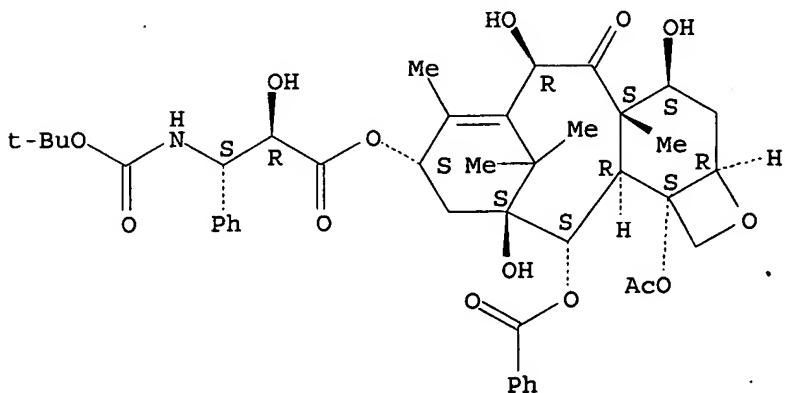
IT 114977-28-5, Docetaxel

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(fused imidazole derivs., and preparation thereof, for improving oral bioavailability of pharmaceutical agents)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

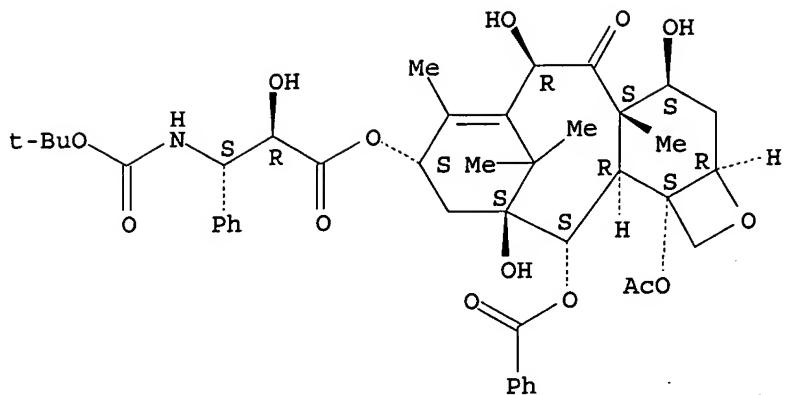


TITLE: Down-regulation of DNA repair to enhance sensitivity
 to p53-mediated suppression in cancer therapy
 INVENTOR(S): Gjerset, Ruth A.
 PATENT ASSIGNEE(S): Sidney Kimmel Cancer Center, USA; Gjerset, Ruth A.
 SOURCE: PCT Int. Appl., 88 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 9801123 | A1 | 19980115 | WO 1997-US12542 | 19970702 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
DK, EE, ES, FI, GB, GE, GH, HU, IL, IS, JP, KE, KG, KP, KR, KZ,
LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL,
PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US,
UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR,
GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA,
GN, ML, MR, NE, SN, TD, TG | | | | |
| US 6054467 | A | 20000425 | US 1996-675887 | 19960705 <-- |
| CA 2259960 | AA | 19980115 | CA 1997-2259960 | 19970702 <-- |
| AU 9736705 | A1 | 19980202 | AU 1997-36705 | 19970702 <-- |
| AU 724212 | B2 | 20000914 | | |
| EP 910357 | A1 | 19990428 | EP 1997-933543 | 19970702 <-- |
| EP 910357 | B1 | 20030604 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI | | | | |
| JP 2000516207 | T2 | 20001205 | JP 1998-505400 | 19970702 <-- |
| AT 241973 | E | 20030615 | AT 1997-933543 | 19970702 |
| US 2005095226 | A1 | 20050505 | US 2004-842718 | 20040510 |
| PRIORITY APPLN. INFO.: | | | US 1996-675887 | A2 19960705 |
| | | | WO 1997-US12542 | W 19970702 |
| | | | US 2000-556440 | B1 20000424 |

- AB The present invention details methods for the treatment of cancer. In particular, it concerns the induction of apoptosis in cancer cells following treatment with inhibitors of DNA repair in combination with p53 gene therapy. Treatment of glioblastoma and breast tumor cells with inhibitors of DNA repair induced growth suppression that was a result of p53-mediated apoptosis. Thus it appears that inhibitors of DNA repair in combination with p53 gene therapy is involved in restoration of p53-mediated apoptosis.
- IT 114977-28-5, Taxotere
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (down-regulation of DNA repair to enhance sensitivity to p53-mediated suppression in cancer therapy)
- RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 17 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 1999:405000 CAPLUS

DOCUMENT NUMBER: 131:43591

TITLE: Combination therapy of cancer with anti-ErbB2 antibodies

INVENTOR(S): Shak, Steven; Paton, Virginia E.

PATENT ASSIGNEE(S): Genentech, Inc., USA

SOURCE: PCT Int. Appl., 42 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-------------------|--------------|
| WO 9931140 | A1 | 19990624 | WO 1998-US26266 | 19981210 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IS, JP, KE,
KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW,
MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR,
TT, UA, UG, UZ, VN, YU, ZW
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| ZA 9811162 | A | 20000607 | ZA 1998-11162 | 19981207 <-- |
| CA 2311409 | AA | 19990624 | CA 1998-2311409 | 19981210 <-- |
| AU 9919081 | A1 | 19990705 | AU 1999-19081 | 19981210 <-- |
| EP 1037926 | A1 | 20000927 | EP 1998-963840 | 19981210 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO | | | | |
| TR 200001689 | T2 | 20010122 | TR 2000-200001689 | 19981210 |
| CN 1281468 | A | 20010124 | CN 1998-812097 | 19981210 |
| BR 9815363 | A | 20011016 | BR 1998-15363 | 19981210 |
| JP 2002508397 | T2 | 20020319 | JP 2000-539062 | 19981210 |
| CN 1820734 | A | 20060823 | CN 2006-10008639 | 19981210 |
| NZ 504597 | A | 20030530 | NZ 2000-504597 | 20000517 |
| NO 2000002957 | A | 20000811 | NO 2000-2957 | 20000609 <-- |
| US 2003147884 | A1 | 20030807 | US 2003-356824 | 20030203 |
| US 2004037823 | A9 | 20040226 | | |
| US 2003170234 | A1 | 20030911 | US 2003-406925 | 20030404 |
| US 2005002928 | A1 | 20050106 | US 2004-909998 | 20040802 |
| PRIORITY APPLN. INFO.: | | | US 1997-69346P | P 19971212 |
| | | | CN 1998-812097 | A3 19981210 |
| | | | US 1998-208649 | A3 19981210 |

US 1998-209023 A3 19981210
WO 1998-US26266 W 19981210

AB The authors disclose the treatment of disorders characterized by the overexpression of ErbB2. More specifically, human patients are treated with a combination of an anti-ErbB2 antibody and a chemotherapeutic agent other than an anthracycline (e.g., doxorubicin or epirubicin). Preferably, the chemotherapeutic agent is Taxol.

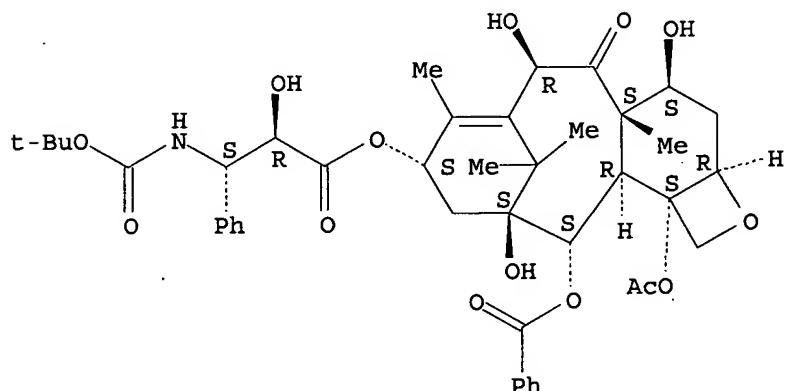
IT 114977-28-5

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(in combination cancer therapy with anti-erbB-2 receptor antibodies)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 6 THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 18 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:911036 CAPLUS

DOCUMENT NUMBER: 134:76383

TITLE: Oral pharmaceutical compositions containing taxanes

INVENTOR(S): Gutierrez-Rocca, Jose C.; Cacace, Janice L.; Selim, Sami; Testman, Robert; Rutledge, J. Michael

PATENT ASSIGNEE(S): Baker Norton Pharmaceuticals, Inc., USA

SOURCE: PCT Int. Appl., 48 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---------------|--|----------|-----------------|-------------|
| WO 2000078247 | A1 | 20001228 | WO 1999-US13821 | 19990618 <- |
| W: | AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZA, ZW | | | |
| RW: | GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |

| | | | |
|--|-------------|-----------------|--------------|
| CA 2371924 | AA 20001228 | CA 1999-2371924 | 19990618 <-- |
| AU 9946955 | A1 20010109 | AU 1999-46955 | 19990618 |
| AU 774060 | B2 20040617 | | |
| BR 9917403 | A 20020709 | BR 1999-17403 | 19990618 |
| EP 1221908 | A1 20020717 | EP 1999-930408 | 19990618 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI, RO, CY | | | |
| JP 2003502349 | T2 20030121 | JP 2001-504316 | 19990618 |
| HU 200300836 | A2 20030828 | HU 2003-836 | 19990618 |
| NZ 516279 | A 20040625 | NZ 1999-516279 | 19990618 |
| RU 2236226 | C2 20040920 | RU 2002-100703 | 19990618 |
| EP 1479382 | A1 20041124 | EP 2004-77062 | 19990618 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI, RO, CY | | | |
| PRIORITY APPLN. INFO.: | | EP 1999-930408 | A3 19990618 |
| | | WO 1999-US13821 | W 19990618 |

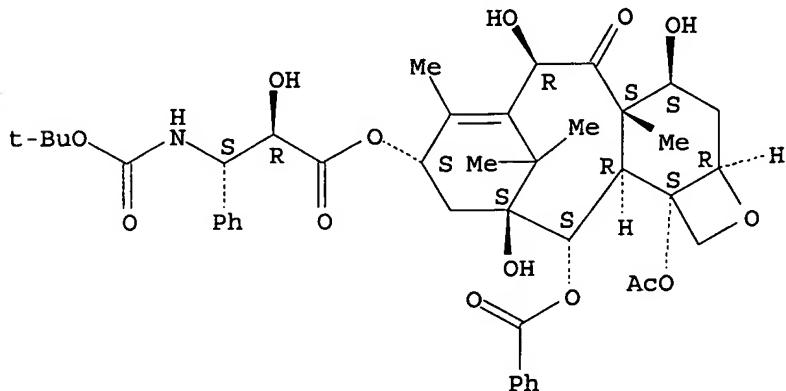
AB Pharmaceutical compns. for oral administration to mammalian subjects comprise a taxane or taxane derivative (e.g., paclitaxel or docetaxel) as active ingredient and a vehicle comprising at least 30% by weight of a carrier for the taxane, the carrier having an HLB value of at least about 10. The compns. may also comprise 0-70% of a viscosity-reducing co-solubilizer. The compns. may be incorporated into conventional oral pharmaceutical dosage forms, or can be in the form of a 2-part drug wherein the first part includes the taxane in a solubilizing vehicle and the second part comprises a carrier for the taxane to promote oral absorption. Methods of treatment of taxane-responsive disease conditions employing the novel compns. are also disclosed, whereby the compns. can be administered alone or in association with an oral bioavailability enhancing agent. A formulation containing Tween 80 at 18 mg/kg and paclitaxel gave an absolute bioavailability of 54% which was >15% for i.v. drug.

IT 114977-28-5, Docetaxel
 RL: BPR (Biological process); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); PROC (Process); USES (Uses)
 (oral pharmaceuticals containing taxanes)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD, ALL CITATIONS AVAILABLE IN THE RE FORMAT

ACCESSION NUMBER: 2005:1210172 CAPLUS
 DOCUMENT NUMBER: 143:466194
 TITLE: Oral pharmaceutical compositions containing taxanes and methods of cancer therapy employing the same
 INVENTOR(S): Gutierrez-Rocca, Jose C.; Cacace, Janice L.; Selim, Sami; Testman, Robert; Rutledge, J. Michael
 PATENT ASSIGNEE(S): Baker Norton Pharmaceuticals, Inc., USA
 SOURCE: U.S., 14 pp., Cont.-in-part of U.S. Ser. No. 863,513, abandoned.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 4
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-----------------|--------------|
| US 6964946 | B1 | 20051115 | US 1998-55818 | 19980406 |
| US 5968972 | A | 19991019 | US 1996-608776 | 19960229 <-- |
| US 6245805 | B1 | 20010612 | US 1996-733142 | 19961016 |
| ZA 9609001 | A | 19970617 | ZA 1996-9001 | 19961025 <-- |
| NZ 516026 | A | 20030630 | NZ 1998-516026 | 19980422 |
| AU 2002035584 | A5 | 20020606 | AU 2002-35584 | 20020422 |
| AU 784159 | B2 | 20060216 | | |
| US 2005267201 | A1 | 20051201 | US 2005-165896 | 20050624 |
| PRIORITY APPLN. INFO.: | | | | |
| | | | US 1995-7071P | P 19951026 |
| | | | US 1996-608776 | A2 19960229 |
| | | | US 1996-733142 | A2 19961016 |
| | | | US 1997-863513 | B2 19970527 |
| | | | US 1998-55818 | A3 19980406 |
| | | | AU 1998-71300 | A3 19980422 |
| | | | NZ 1998-501127 | A1 19980422 |

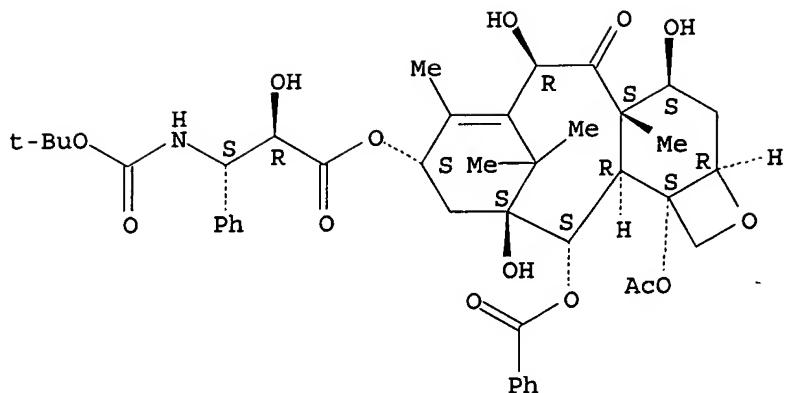
AB The present invention relates to pharmaceutical compns. for oral administration to mammalian subjects comprising a taxane or taxane derivative (e.g., paclitaxel or docetaxel) as active ingredient and a vehicle comprising at least 30% by weight of a carrier for the taxane, said carrier having an HLB value of at least about 10. The compns. may also comprise 0-70% of a viscosity-reducing co-solubilizer. The compns. may be incorporated into conventional oral pharmaceutical dosage forms, or can be in the form of a two-part medicament wherein the first part includes the taxane in a solubilizing vehicle and the second part comprises a carrier for the taxane to promote oral absorption. Methods of treatment of taxane-responsive disease conditions employing the novel compns. are also disclosed, whereby the compns. can be administered alone or in association with an oral bioavailability enhancing agent.

IT 114977-28-5, Docetaxel
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (oral pharmaceutical compns. containing taxanes and methods of cancer therapy employing same)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 20 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:191189 CAPLUS..

DOCUMENT NUMBER: 132:227475

TITLE: Treatment of oncologic tumors with an injectable formulation of a Golgi apparatus disturbing agent

INVENTOR(S): Singh, Saira Sayed

PATENT ASSIGNEE(S): Oncopharmaceutical, Inc., USA

SOURCE: PCT Int. Appl., 32 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|------|----------|-----------------|--------------|
| WO 2000015766 | A1 | 20000323 | WO 1999-US21312 | 19990915 <-- |
| W: AU, CA, JP, KR
RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
PT, SE | | | | |
| CA 2344316 | AA | 20000323 | CA 1999-2344316 | 19990915 <-- |
| AU 9959253 | A1 | 20000403 | AU 1999-59253 | 19990915 <-- |
| EP 1114144 | A1 | 20010711 | EP 1999-946955 | 19990915 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI | | | | |
| US 6287602 | B1 | 20010911 | US 1999-397390 | 19990915 |
| JP 2002525268 | T2 | 20020813 | JP 2000-570293 | 19990915 |
| US 2002012703 | A1 | 20020131 | US 2001-912115 | 20010723 |
| US 6497904 | B2 | 20021224 | | |
| PRIORITY APPLN. INFO.: | | | US 1998-100479P | P 19980916 |
| | | | US 1999-397390 | A1 19990915 |
| | | | WO 1999-US21312 | W 19990915 |

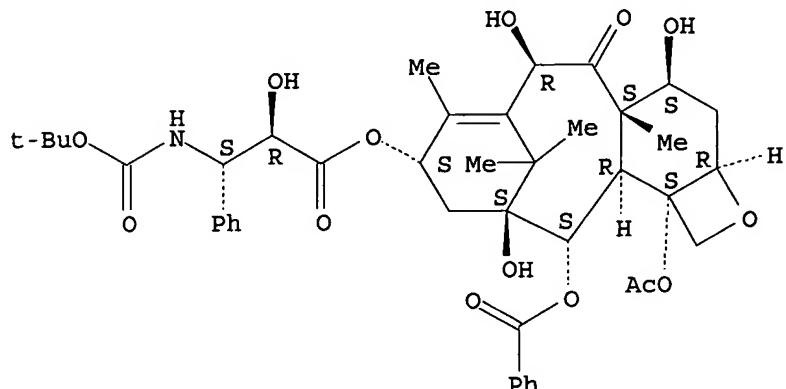
AB Novel pharmaceutical formulations for treating a cellular proliferative disease are provided comprising: a therapeutically effective amount of a Golgi apparatus disturbing agent; a biocompatible carrier; and a solvent. In preferred formulations, the Golgi apparatus disturbing agent is brefeldin A (BFA) and the biocompatible carrier is a polymer such as chitin or chitosan. Methods of treating cellular proliferative diseases using the pharmaceutical formulations are also described. Nude mice bearing human epithelial (KB-1) tumors were treated with a BFA/chitin/dimethylacetamide composition

IT 114977-28-5, Docetaxel

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(as addnl. pharmacol. agent; treatment of oncol. tumors with injectable formulation of golgi apparatus disturbing agent).

RN 114977-28-5 CAPLUS
CN Benzenepropanoic acid, β -[(1,1-dimethylethoxy)carbonyl]amino]-
 α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-
(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-
trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-
cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA
INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 21 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2000:456819 CAPLUS
DOCUMENT NUMBER: 133:84238
TITLE: 3-heteroarylidenyl-2-indolinone compounds for modulating protein kinase activity and for use in cancer chemotherapy
INVENTOR(S): Langecker, Peter J.; Shawver, Laura Kay; Tang, Peng Cho; Sun, Li
PATENT ASSIGNEE(S): Sugen, Inc., USA
SOURCE: PCT Int. Appl., 148 pp.
CODEN: PIXXD2
DOCUMENT TYPE: Patent
LANGUAGE: English
FAMILY ACC. NUM. COUNT: 2
PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 2000038519 | A1 | 20000706 | WO 1999-US31232 | 19991230 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2357042 | AA | 20000706 | CA 1999-2357042 | 19991230 <-- |
| BR 9916735 | A | 20010925 | BR 1999-16735 | 19991230 |
| EP 1139754 | A1 | 20011010 | EP 1999-966725 | 19991230 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| JP 2002533360 | T2 | 20021008 | JP 2000-590484 | 19991230 |
| AU 760964 | B2 | 20030522 | AU 2000-22215 | 19991230 |
| WO 2001049287 | A1 | 20010712 | WO 2000-US18058 | 20000630 |

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 CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
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 LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
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 DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
 CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG
 EP 1259234 A1 20021127 EP 2000-943334 20000630
 EP 1259234 B1 20060816
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, SI, LT, LV, FI, RO, MK, CY, AL
 JP 2003535038 T2 20031125 JP 2001-549655 20000630
 AT 336245 E 20060915 AT 2000-943334 20000630
 US 2003191162 A1 20031009 US 2002-307483 20021202
 PRIORITY APPLN. INFO.: US 1998-114313P P 19981231
 US 1999-476232 A 19991230
 WO 1999-US31232 W 19991230
 US 2000-569545 A 20000512
 WO 2000-US18058 W 20000630

OTHER SOURCE(S): MARPAT 133:84238

AB 3-Heteroarylidene-2-indolinone compds. are provided that modulate the enzymic activity of protein kinases and therefore are expected to be useful in the prevention and treatment of protein kinase-related cellular disorders, e.g. cancer. Furthermore, these compds. are expected to enhance the efficacy of other chemotherapeutic agents, in particular, fluorinated pyrimidines, in the treatment of cancer.

IT 114977-28-5, Docetaxel

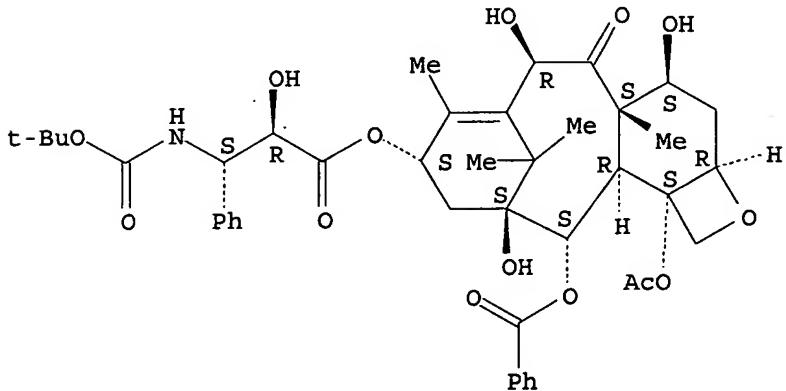
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(heteroarylideneindolinone derivs. for modulating protein kinase activity and in cancer chemotherapy)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

DOCUMENT NUMBER: 134:37055
 TITLE: Methods and compositions using FGF inhibitors and
 agonists for modulating cell proliferation and cell
 death
 INVENTOR(S): Au, Jessie L. S.; Wientjes, M. Guillaume
 PATENT ASSIGNEE(S): USA
 SOURCE: PCT Int. Appl., 143 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|-----------------|-----------------|--------------|
| WO 2000074634 | A2 | 20001214 | WO 2000-US40103 | 20000605 <-- |
| WO 2000074634 | C2 | 20020926 | | |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU,
CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID,
IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV,
MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, PL, PT, RO, RU, SD, SE,
SG, SI, SK
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY,
DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ,
CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2377385 | AA | 20001214 | CA 2000-2377385 | 20000605 <-- |
| EP 1206234 | A2 | 20020522 | EP 2000-943429 | 20000605 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL | | | | |
| JP 2003503313 | T2 | 20030128 | JP 2001-501171 | 20000605 |
| US 6599912 | B1 | 20030729 | US 2000-587559 | 20000605 |
| AU 780454 | B2 | 20050324 | AU 2000-57903 | 20000605 |
| US 2004010001 | A1 | 20040115 | US 2003-464018 | 20030618 |
| PRIORITY APPLN. INFO.: | | | | |
| | | US 1999-137345P | P | 19990603 |
| | | US 1999-165983P | P | 19991117 |
| | | US 1999-172031P | P | 19991223 |
| | | US 2000-187445P | P | 20000307 |
| | | US 2000-587559 | A3 | 20000605 |
| | | WO 2000-US40103 | W | 20000605 |

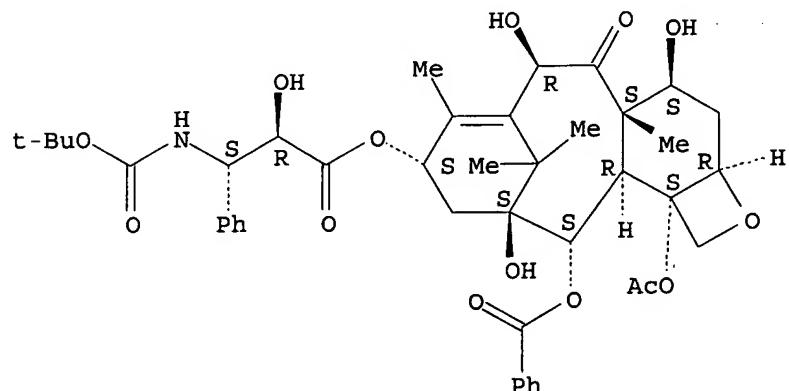
- AB Methods and compns. for modulating the FGF effect on the sensitivity of malignant and normal cells to anticancer agents are provided. In particular, methods and compns. for inhibiting FGF-induced resistance to a broad spectrum of anticancer agents in solid and soft-tissue tumors, metastatic lesions, leukemia and lymphoma are provided. Preferably, the compns. include at least one FGF inhibitor in combination with a cytotoxic agents, e.g., antimicrotubule agents, topoisomerase I inhibitors, topoisomerase II inhibitors, antimetabolites, mitotic inhibitors, alkylating agents, intercalating agents, agents capable of interfering with a signal transduction pathway (e.g., g., a protein kinase C inhibitor, e.g., an anti-hormone, e.g., an antibody against growth factor receptors), an agent that promote apoptosis and/or necrosis, an interferon, an interleukin, a tumor necrosis factor, and radiation. In other embodiments, methods and composition for protecting a cell in a subject, from one or more of killing, inhibition of growth or division or other damage caused, e.g., by a cytotoxic agent, are provided. Preferably, the method includes administering to the subject an effective amount of at least one FGF agonist, thereby treating the cell, e.g., protecting or reducing the damage to the dividing cell from said cytotoxic agent. FGF gene expression-based methods for diagnosis of proliferative disorders are also disclosed.
- IT 114977-28-5, Taxotere
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(FGF inhibitors and agonists for modulating cell proliferation and cell death)

RN 114977-28-5 CAPIUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 23 OF 55 CAPIUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2000:824125 CAPIUS

DOCUMENT NUMBER: 134:4050

TITLE: Treatment with anti-erbB2 antibodies

INVENTOR(S): Cohen, Robert L.

PATENT ASSIGNEE(S): Genentech, Inc., USA

SOURCE: PCT Int. Appl., 39 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

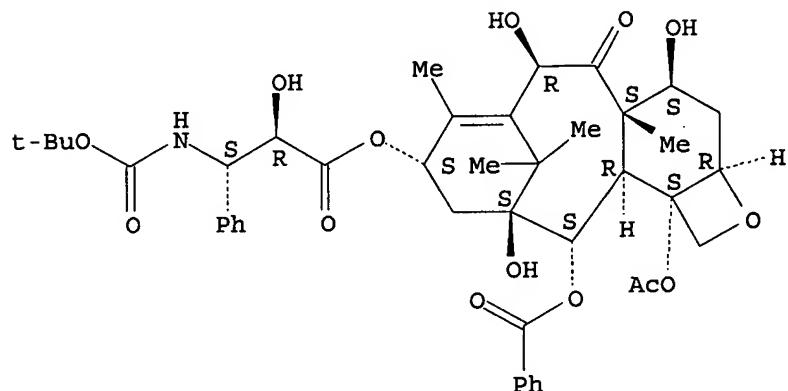
FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 2000069460 | A1 | 20001123 | WO 2000-US12552 | 20000509 <-- |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2374085 | AA | 20001123 | CA 2000-2374085 | 20000509 <-- |
| EP 1187632 | A1 | 20020320 | EP 2000-928916 | 20000509 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO | | | | |
| JP 2002544238 | T2 | 20021224 | JP 2000-617920 | 20000509 |
| AU 782325 | B2 | 20050721 | AU 2000-47080 | 20000509 |
| US 2003170235 | A1 | 20030911 | US 2003-429519 | 20030505 |
| PRIORITY APPLN. INFO.: | | | US 1999-134085P | P 19990514 |
| | | | US 2000-568322 | A1 20000509 |
| | | | WO 2000-US12552 | W 20000509 |

AB A method treating a human patient to or diagnosed with a tumor in which erbB2 protein is expressed comprising the following steps, performed sequentially: (a) treating the patient with a therapeutically effective amount of an anti-erbB2 antibody; (b) surgically removing the tumor, and then (c) treating the patient with a therapeutically effective amount of an anti-erbB2 antibody or of a chemotherapeutic agent.
 IT 114977-28-5
 RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (cancer treatment with anti-erbB2 antibodies)
 RN 114977-28-5 CAPLUS
 CN Benzene propanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]-
 α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-
 (benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-
 trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-
 cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA
 INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 24 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2002:158387 CAPLUS
 DOCUMENT NUMBER: 136:210551
 TITLE: Method of treating hyperproliferative diseases using active vitamin D analogues
 INVENTOR(S): Bishop, Charles W.; Mazess, Richard B.
 PATENT ASSIGNEE(S): Bone Care International, Inc., USA
 SOURCE: U.S. Pat. Appl. Publ., 13 pp., Cont.-in-part of U.S. Ser. No. 596,149.
 CODEN: USXXCO
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 20
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---------------|------|----------|-----------------|--------------|
| US 2002025950 | A1 | 20020228 | US 2001-891814 | 20010626 |
| US 6503893 | B2 | 20030107 | | |
| US 5763429 | A | 19980609 | US 1996-781910 | 19961230 <-- |
| US 6537982 | B1 | 20030325 | US 1998-596149 | 19980223 |
| US 2002128240 | A1 | 20020912 | US 2001-995911 | 20011128 |
| CA 2450942 | AA | 20030103 | CA 2002-2450942 | 20020626 |
| WO 2003000023 | A2 | 20030103 | WO 2002-US20475 | 20020626 |
| WO 2003000023 | A3 | 20030731 | | |

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,

CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
 GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
 LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
 PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
 UA, UG, UZ, VN, YU, ZA, ZM, ZW
 RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
 KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB,
 GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA,
 GN, GQ, GW, ML, MR, NE, SN, TD, TG
 EP 1408983 A2 20040421 EP 2002-756332 20020626
 R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
 IE, FI, CY, TR
 CN 1520302 A 20040811 CN 2002-812881 20020626
 JP 2004535429 T2 20041125 JP 2003-506479 20020626
 US 2003130242 A1 20030710 US 2003-337506 20030107
 US 6680309 B2 20040120
 PRIORITY APPLN. INFO.: US 1996-781910 A3 19961230
 US 1998-596149 A2 19980223
 US 1993-119895 A2 19930910
 US 1994-265438 A2 19940624
 US 1995-415488 A2 19950403
 US 1995-486387 A2 19950607
 US 2001-891814 A2 20010626
 WO 2002-US20475 W 20020626

OTHER SOURCE(S): MARPAT 136:210551

AB Methods use hypocalcemic vitamin D analogs to inhibit the hyperproliferation of malignant or neoplastic cells without incidence of hypercalcemia. Patients with advanced androgen-independent prostate cancer were treated with 1 α ,24-dihydroxyvitamin D2.

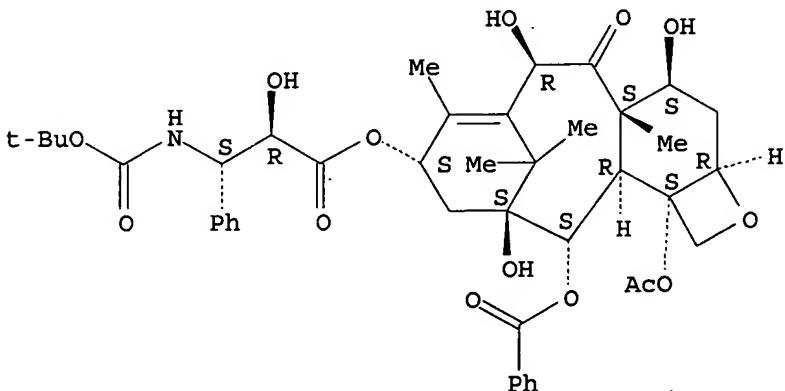
IT 114977-28-5, Docetaxel

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (coadministration with cytotoxic; hypocalcemic vitamin D analogs for treating hyperproliferative diseases)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



TITLE: Treatment of patients with liver metastases
 AUTHOR(S): Fumoleau, P.
 CORPORATE SOURCE: Center Regionale de Lutte Contre le Cancer,
 Nantes-Atlantique, Herblain, 44805, Fr.
 SOURCE: Anti-Cancer Drugs (1996), 7(Suppl. 2,
 Management of Advanced Breast Cancer: Patient Needs,
 Challenges and New Treatment Options), 21-23.
 CODEN: ANTDEV; ISSN: 0959-4973
 PUBLISHER: Rapid Science Publishers
 DOCUMENT TYPE: Journal
 LANGUAGE: English

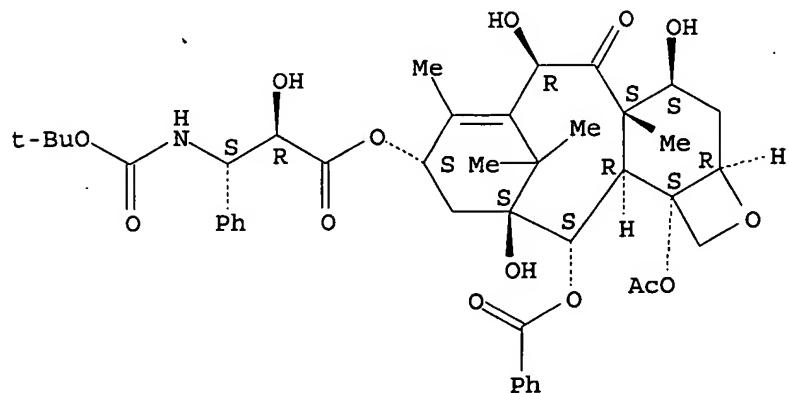
AB The presence of liver metastases is a very poor prognostic factor for patients with metastatic breast cancer. Liver metastases are generally less responsive to chemotherapy than metastases in other sites, and patients with liver lesions have a shorter survival duration than patients with other sites of disease. The results from 5 multicenter phase II studies of docetaxel as a first-line treatment for metastatic breast cancer were analyzed with regard to the presence or absence of liver lesions, which were found in 39% of the 209 patients involved. Response rates to docetaxel, 100 or 75 mg/m², were maintained in the presence of liver lesions and the median survival across all five studies was 16.4 mo for all patients and 14.7 mo for patients with liver lesions. Similarly, when results from 129 patients given docetaxel as a second-line treatment were analyzed, the response rates and survival durations were not reduced in the 57% of patients who had liver lesions. The presence of liver metastases does not reduce the probability or duration of response to docetaxel as a first- or second-line treatment for advanced breast cancer.

IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (treatment of patients with liver metastases)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 26 OF 55 MEDLINE on STN
 ACCESSION NUMBER: 2000024224 MEDLINE
 DOCUMENT NUMBER: PubMed ID: 10560434
 TITLE:

A case of multiple liver metastases from breast cancer

successfully treated with intra-arterial administration of docetaxel.
 AUTHOR: Maeda Y; Nishida M; Takao T; Harada K; Mori N; Tamesa T;
 Somura H; Tangoku A; Oka M; Konishi T
 CORPORATE SOURCE: Dept. of Surgery II, Yamaguchi University School of Medicine.
 SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1999 Oct) Vol. 26, No. 12, pp. 1951-4.
 Journal code: 7810034. ISSN: 0385-0684.
 PUB. COUNTRY: Japan
 DOCUMENT TYPE: (CASE REPORTS)
 Journal; Article; (JOURNAL ARTICLE)
 LANGUAGE: Japanese
 FILE SEGMENT: Priority Journals
 ENTRY MONTH: 199911
 ENTRY DATE: Entered STN: 13 Jan 2000
 Last Updated on STN: 13 Jan 2000
 Entered Medline: 26 Nov 1999

AB Docetaxel is an excellent agent with a high antitumor effect for the treatment of advanced/recurrent breast cancer. A 55-year-old female with metastatic liver tumors from breast cancer showed a remarkable response to intra-arterial administration of docetaxel (20 mg/week, or 40 mg/2 weeks). Since CT and MRI imaging revealed multiple metastases in the liver, intra-arterial chemotherapy was selected. No critical side effect was found during this chemotherapy. A CT scan 3 months after chemotherapy showed a partial response. We conclude that this intra-arterial chemotherapy using docetaxel will be safe and useful for liver metastases from breast cancer.

L12 ANSWER 27 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1998:564267 CAPLUS
 DOCUMENT NUMBER: 129:197984
 TITLE: Combined tumor suppressor gene therapy and chemotherapy in the treatment of neoplasms
 INVENTOR(S): Nielsen, Loretta; Horowitz, Jo Ann; Maneval, Daniel C.; Demers, G. William; Rybak, Mary Ellen; Resnick, Gene
 PATENT ASSIGNEE(S): Canji, Inc., USA
 SOURCE: PCT Int. Appl., 114 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 9835554 | A2 | 19980820 | WO 1998-US3514 | 19980217 <-- |
| WO 9835554 | A3 | 19981126 | | |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GE, GH, GM, GW, HU, ID, IL, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, US, UZ, VN, YU, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, ML, MR, NE, SN, TD, TG | | | | |
| CA 2282683 | AA | 19980820 | CA 1998-2282683 | 19980217 <-- |
| AU 9864380 | A1 | 19980908 | AU 1998-64380 | 19980217 <-- |
| AU 737621 | B2 | 20010823 | | |
| EP 969720 | A2 | 20000112 | EP 1998-910038 | 19980217 <-- |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, FI | | | | |

| | | | | |
|------------------------|----|----------------|----------------|--------------|
| NZ 337283 | A | 20010223 | NZ 1998-337283 | 19980217 |
| HU 200004326 | A2 | 20010228 | HU 2000-4326 | 19980217 |
| JP 2001511815 | T2 | 20010814 | JP 1998-536033 | 19980217 |
| BR 9807418 | A | 20020122 | BR 1998-7418 | 19980217 |
| US 2003060434 | A1 | 20030327 | US 1999-311772 | 19990513 |
| NO 9903943 | A | 19991015 | NO 1999-3943 | 19990817 <-- |
| US 2003064949 | A1 | 20030403 | US 2002-86294 | 20020228 |
| US 2004235736 | A1 | 20041125 | US 2004-824058 | 20040413 |
| US 2005142112 | A1 | 20050630 | US 2004-823932 | 20040413 |
| PRIORITY APPLN. INFO.: | | | | |
| | | US 1997-38065P | P 19970218 | |
| | | US 1997-801285 | A 19970218 | |
| | | US 1997-801681 | A 19970218 | |
| | | US 1997-801755 | A 19970218 | |
| | | US 1997-801765 | A 19970218 | |
| | | US 1997-47834P | P 19970528 | |
| | | US 1998-24932 | B1 19980217 | |
| | | WO 1998-US3514 | W 19980217 | |
| | | US 1999-311772 | B3 19990513 | |

AB In one embodiment, the invention provides methods of treating mammalian cancer or hyperproliferative cells, the method comprising contacting the cells with a tumor suppressor protein or tumor suppressor nucleic acid and also contacting the cells with at least one adjunctive anticancer agent. The invention also provides for a pharmacol. composition comprising a tumor suppressor protein or a tumor suppressor nucleic acid and at least one adjunctive anti-cancer agent, as well as a kit for the treatment of mammalian cancer or hyperproliferative cells.

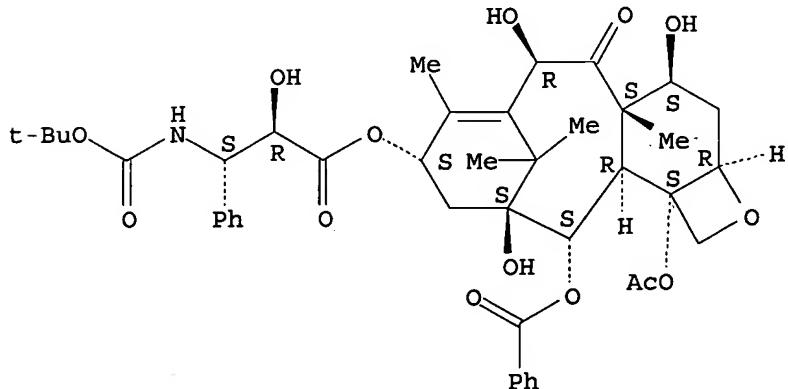
IT 114977-28-5, Taxotere
RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tumor suppressor gene therapy-chemotherapy combination for treatment of neoplasms and hyperproliferative cells)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -{[(1,1-dimethylethoxy)carbonyl]amino}- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 28 OF 55 MEDLINE on STN

ACCESSION NUMBER: 2001066728 MEDLINE

DOCUMENT NUMBER: PubMed ID: 10907946

TITLE: Phase II study of docetaxel in patients with liver metastases from breast cancer. UK study group.

AUTHOR: Coleman R E; Howell A; Eggleton S P; Maling S J; Miles D W
CORPORATE SOURCE: Weston Park Hospital NHS Trust, Sheffield, UK.
SOURCE: Annals of oncology : official journal of the European Society for Medical Oncology / ESMO, (2000 May)
Vol. 11, No. 5, pp. 541-6.
Journal code: 9007735, ISSN: 0923-7534.

PUB. COUNTRY: Netherlands
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
(MULTICENTER STUDY)

LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200012
ENTRY DATE: Entered STN: 22 Mar 2001
Last Updated on STN: 22 Mar 2001
Entered Medline: 28 Dec 2000

AB BACKGROUND: Previous phase II studies of docetaxel have indicated that hepatic metastases from breast cancer respond well to first-line treatment with docetaxel. The objective of this prospective, open label phase II study therefore was specifically to evaluate the activity and safety of docetaxel in this indication. PATIENTS AND METHODS: The study recruited 47 women (mean age 50 years, range 33-66 years) with hepatic metastases from breast cancer who fulfilled the eligibility criteria. After premedication with steroids, patients received a one-hour intravenous infusion of docetaxel 100 mg/m² at three-weekly intervals for up to eight cycles. Response to treatment during medication was assessed after three, six and where appropriate, eight cycles and every three month follow-up thereafter, until disease progression or death. RESULTS: The best overall response rate (ORR) for evaluable patients was 64.3% (95% CI: 48.0-78.5%). In terms of the primary efficacy parameters, the ORR at the sixth cycle of treatment was 62% (95% CI: 45%-80%) with 17% complete responses. The median duration of response was 139 days (95% CI: 111-216 days) and the median survival duration calculated on an intent-to-treat basis was 335 days (227-568 days, 95% CI). One (2%) toxic death was reported. CONCLUSIONS: Docetaxel is a highly effective cytotoxic agent in the treatment of patients with liver metastases from breast cancer.

L12 ANSWER 29 OF 55 MEDLINE on STN
ACCESSION NUMBER: 1998056505 MEDLINE
DOCUMENT NUMBER: PubMed ID: 9446016
TITLE: [Docetaxel (taxotere) for therapy of breast carcinoma. Highest effectiveness with moderate side effects].
Docetaxel (Taxotere) zur Therapie des Mammakarzinoms. Hochste Wirksamkeit bei moderaten Nebenwirkungen.
AUTHOR: von Minckwitz G; Costa S D
CORPORATE SOURCE: Klinik fur Gynakologie und Geburtshilfe, Johann Wolfgang Goethe-Universitat Frankfurt.. minckwitz@em.uni-frankfurt.de
SOURCE: Medizinische Klinik (Munich, Germany : 1983), (1997 Sep 15) Vol. 92 Suppl 4, pp. 4-9. Ref: 16
Journal code: 8303501. ISSN: 0723-5003.
PUB. COUNTRY: GERMANY: Germany, Federal Republic of
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)
General Review; (REVIEW)
LANGUAGE: German
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199801
ENTRY DATE: Entered STN: 6 Feb 1998
Last Updated on STN: 6 Feb 1998
Entered Medline: 27 Jan 1998

AB CLINICAL RESULTS: Docetaxel is a taxan which has proven high efficacy in the treatment of breast cancer. The results are consistent throughout all phases of clinical evaluation. High response rates have been observed especially for women after failure of anthracyclins or with liver metastases. Response rates are superior to doxorubicin, while the extent of the side effects is comparable. CONCLUSION: Due to the different toxicity profile a combination of docetaxel and anthracyclins is feasible and has already been demonstrated in early clinical trials. The role of the combinatory treatments in first line or adjuvant setting is currently under investigation.

L12 ANSWER 30 OF 55 MEDLINE on STN
ACCESSION NUMBER: 96351131 MEDLINE
DOCUMENT NUMBER: PubMed ID: 8745348
TITLE: Docetaxel: a new defence in the management of breast cancer.
AUTHOR: Piccart M
CORPORATE SOURCE: Department of Chemotherapy, Institut Jules Bordet, Brussels, Belgium.
SOURCE: Anti-cancer drugs, (1995 Jul) Vol. 6 Suppl 4, pp. 7-11. Ref: 12.
PUB. COUNTRY: ENGLAND: United Kingdom
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)
General Review; (REVIEW)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199610
ENTRY DATE: Entered STN: 25 Oct 1996
Last Updated on STN: 25 Oct 1996
Entered Medline: 16 Oct 1996

AB The results of nine phase II trials of docetaxel in the first- and second-line treatment of patients with advanced breast cancer are summarized. All 316 patients included in this report received docetaxel at a dose of 100 mg/m² administered over 1 h every 3 weeks on an outpatient basis. One hundred and fifty-four patients received docetaxel as first-line therapy for advanced disease, half of whom had received prior adjuvant chemotherapy (finished at least 1 year previously). An overall response rate of 59% (95% CI: 51-67) was achieved in these patients, with a median duration of response of 8.3 months and a median time to progression of 4.9 months. Similar results were seen in a subgroup of 68 patients with liver metastases. Among the 162 patients given docetaxel as second-line therapy, 134 had strictly defined anthracycline-resistant disease; 73 had liver metastases. The combined overall response rate for anthracycline-resistant patients in two US studies was 48% (95% CI: 37-59) while that in a multicenter French study was 29% (95% CI: 18-44). The median duration of response in each case was 6.3 and 5.5 months, respectively, with an overall median survival duration of 11 and 10 months, respectively. Among patients with liver metastases, second-line treatment with docetaxel achieved an overall response rate of 32%, a median duration of response of 7.8 months and a median survival duration of 9 months. These results for docetaxel as both first- and second-line therapy are comparable with those achieved with doxorubicin and are particularly promising in patients with liver metastases and anthracycline-resistant disease.

L12 ANSWER 31 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 1999:736476 CAPLUS
DOCUMENT NUMBER: 131:346535
TITLE: Use of neomycin for treating angiogenesis-related diseases
INVENTOR(S): Hu, Guo-Fu; Vallee, Bert L.
PATENT ASSIGNEE(S): The Endowment for Research In Human Biology, Inc., USA
SOURCE: PCT Int. Appl., 74 pp.

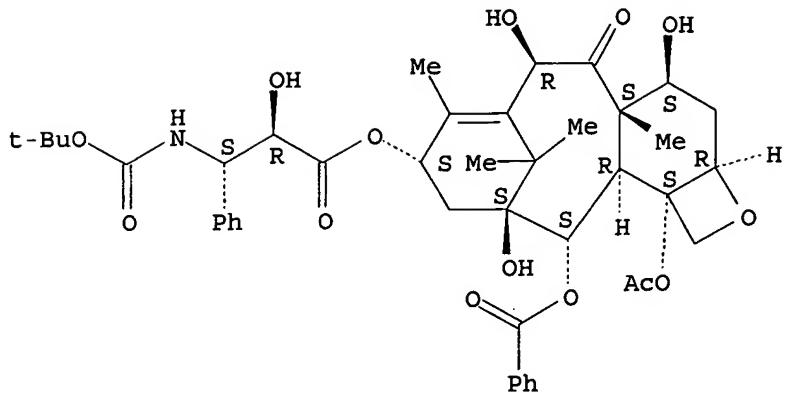
CODEN: PIXXD2

DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|--|--|----------|-----------------|--------------|
| WO 9958126 | A1 | 19991118 | WO 1999-US10269 | 19990511 <-- |
| W: AE, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ,
DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS,
JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK,
MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ,
TM, TR, TT, UA, UG, US, UZ, VN, YU, ZA, ZW, AM, AZ, BY, KG, KZ,
MD, RU, TJ, TM | RW: GH, GM, KE, LS, MW, SD, SL, SZ, UG, ZW, AT, BE, CH, CY, DE, DK,
ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG,
CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | |
| CA 2331620 | AA | 19991118 | CA 1999-2331620 | 19990511 <-- |
| AU 9939804 | A1 | 19991129 | AU 1999-39804 | 19990511 <-- |
| EP 1083896 | A1 | 20010321 | EP. 1999-922915 | 19990511 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, FI | | | | |
| US 6482802 | B1 | 20021119 | US 2000-700436 | 20001109 |
| PRIORITY APPLN. INFO.: | | | US 1998-84921P | P 19980511 |
| | | | WO 1999-US10269 | W 19990511 |

- AB The present invention is directed to using neomycin or an analog thereof as a therapeutic agent to treat angiogenesis-related diseases, which are characterized by excessive, undesired or inappropriate angiogenesis or proliferation of endothelial cells. The present invention is also directed to pharmaceutical compns. comprising: (a) neomycin or an analog and, optionally, (b) another anti-angiogenic agent or an anti-neoplastic agent. The present invention is further directed to a method for screening neomycin analogs having anti-angiogenic activity. A preferred embodiment of the invention relates to using neomycin to treat subjects having such diseases. A dose of 20 ng neomycin/embryo or higher completely inhibited angiogenin-induced angiogenesis in the chorioallantoic membrane (CAM) assay. Neomycin inhibits angiogenin-induced angiogenesis mainly through inhibition of nuclear translocation of angiogenin.
- IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (neomycin, its analogs and other agents for treatment of angiogenesis-related diseases)
- RN 114977-28-5 CAPLUS
- CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

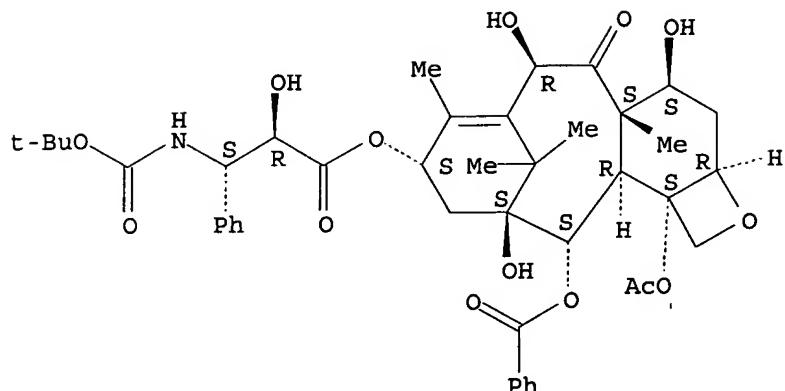
L12 ANSWER 32 OF 55 MEDLINE on STN
 ACCESSION NUMBER: 1999314769 MEDLINE
 DOCUMENT NUMBER: PubMed ID: 10408850
 TITLE: Phase II study of docetaxel in patients with metastatic pancreatic cancer: a Japanese cooperative study. Cooperative Group of Docetaxel for Pancreatic Cancer in Japan.
 AUTHOR: Okada S; Sakata Y; Matsuno S; Kurihara M; Sasaki Y; Ohashi Y; Taguchi T
 CORPORATE SOURCE: Department of Internal Medicine, National Cancer Center Hospital, Tokyo, Japan.
 SOURCE: British journal of cancer, (1999 May) Vol. 80, No. 3-4, pp. 438-43.
 Journal code: 0370635. ISSN: 0007-0920.
 PUB. COUNTRY: SCOTLAND: United Kingdom
 DOCUMENT TYPE: (CLINICAL TRIAL)
 (CLINICAL TRIAL, PHASE II)
 Journal; Article; (JOURNAL ARTICLE)
 (MULTICENTER STUDY)
 LANGUAGE: English
 FILE SEGMENT: Priority Journals
 ENTRY MONTH: 199907
 ENTRY DATE: Entered STN: 27 Jul 1999
 Last Updated on STN: 27 Jul 1999
 Entered Medline: 15 Jul 1999
 AB Docetaxel has been reported to show promising anti-tumour activity in pancreatic ductal cancer (PC). This study was conducted to evaluate the activity and toxicity of moderate-dose (60 mg m⁻²) docetaxel in Japanese chemo-naive patients with measurable metastatic PC. The patients had a performance status of 0-2. They received docetaxel intravenously over a 1- to 2-h period without any premedication for hypersensitivity reactions. This treatment was repeated every 3-4 weeks with dose adjustments based on the toxic effects observed. Twenty-one patients were eligible and treated with docetaxel. The median number of courses was 2 (range, 1-4). None of the patients achieved an objective response; seven showed no change and 13 showed progressive disease. In one patient, the response was not assessable because of early death. The median survival time for all patients was 118 days. The main grade 3-4 toxicities by patient were leucocytopenia (67%) and neutropenia (86%). Other grade 3-4 toxicities included anaemia (10%), thrombocytopenia (5%), nausea/vomiting (29%), anorexia (29%), GOT/GPT increase (10%), alkaline phosphatase increase (14%), malaise/fatigue (33%) and alopecia (24%). In conclusion, docetaxel, administered on this schedule, did not show significant anti-tumour activity in patients with metastatic PC.

L12 ANSWER 33 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 2000:824124 CAPLUS
 DOCUMENT NUMBER: 134:506
 TITLE: Treatment of refractory human tumors with epidermal growth factor receptor antagonists
 INVENTOR(S): Waksal, Harlan W.
 PATENT ASSIGNEE(S): Imclone Systems Incorporated, USA
 SOURCE: PCT Int. Appl., 31 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|-----------------|------------------|--------------|
| WO 2000069459 | A1 | 20001123 | WO 2000-US11756 | 20000501 <-- |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR,
CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU,
ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU,
LV, MA, MD, MG, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE,
SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZW | | | | |
| RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE,
DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF,
CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| CA 2373815 | AA | 20001123 | CA 2000-2373815 | 20000501 <-- |
| BR 2000010524 | A | 20020528 | BR 2000-10524 | 20000501 |
| EP 1218032 | A1 | 20020703 | EP 2000-928671 | 20000501 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
IE, SI, LT, LV, FI, RO, MK, CY, AL | | | | |
| HU 200201480 | A2 | 20020828 | HU 2002-1480 | 20000501 |
| EE 200100603 | A | 20030217 | EE 2001-603 | 20000501 |
| JP 2003520195 | T2 | 20030702 | JP 2000-617919 | 20000501 |
| AU 782994 | B2 | 20050915 | AU 2000-46871 | 20000501 |
| CN 1720994 | A | 20060118 | CN 2005-10055865 | 20000501 |
| US 2002012663 | A1 | 20020131 | US 2001-840146 | 20010424 |
| NO 2001005546 | A | 20020114 | NO 2001-5546 | 20011113 |
| ZA 2001009347 | A | 20030213 | ZA 2001-9347 | 20011113 |
| BG 106110 | A | 20020430 | BG 2001-106110 | 20011114 |
| US 2003157104 | A1 | 20030821 | US 2001-996954 | 20011130 |
| US 2005112120 | A1 | 20050526 | US 2004-18950 | 20041220 |
| PRIORITY APPLN. INFO.: | | | | |
| | | US 1999-312284 | A 19990514 | |
| | | US 1999-374028 | A 19990813 | |
| | | CN 2000-810321 | A3 20000501 | |
| | | WO 2000-US11756 | W 20000501 | |
| | | US 2001-840146 | A1 20010424 | |

- AB A method of inhibiting the growth of refractory tumors that are stimulated by a ligand of epidermal growth factor in human patients comprises treating the human patients with an effective amount of an epidermal growth factor receptor antagonist, e.g. a monoclonal antibody.
 IT 114977-28-5, Docetaxel
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (EGF receptor antagonists for treatment of refractory human tumors)
 RN 114977-28-5 CAPLUS
 CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 3 THERE ARE 3 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 34 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1999:454255 CAPLUS
 DOCUMENT NUMBER: 131:92524
 TITLE: Therapeutic liposome-encapsulated immunomodulators
 INVENTOR(S): Spitler, Lynn E.; Fidler, Issaiyah J.
 PATENT ASSIGNEE(S): Jenner Biotherapies, Inc., USA
 SOURCE: PCT Int. Appl., 111 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 9935162 | A1 | 19990715 | WO 1999-US272 | 19990106 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE,
DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP,
KE, KG, KP, KR, LZ, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN,
MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM,
TR, TT, UA, UG, UZ, VN, YU, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM
RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES,
FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI,
CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 9922141 | A1 | 19990726 | AU 1999-22141 | 19990106 <-- |
| US 2003017976 | A1 | 20030123 | US 2001-764546 | 20010117 |
| US 2004146552 | A1 | 20040729 | US 2003-705618 | 20031110 |
| PRIORITY APPLN. INFO.: | | | US 1998-70717P | P 19980107 |
| | | | US 1999-226075 | B1 19990106 |
| | | | WO 1999-US272 | W 19990106 |
| | | | US 2001-764546 | A1 20010117 |

AB The present invention relates to the use of novel compns. of lipopeptides that are immunomodulators encapsulated as liposomes or free-form for the treatment of neoplasia and in reducing chemotherapeutically induced cellular pathol., including mucositis. These lipopeptides may be administered alone or in combination with a second antineoplastic agent. E.g., a synthetic JBT 3002 lipopeptide entrapped in phosphatidylcholine/phosphatidylserine liposomes is shown to be a potent activator of tumoricidal properties of murine macrophages by a mechanism that differs from that of lipopolysaccharides. These data highly support the in vivo use of multilamellar liposome-encapsulated JBT 3002 to enhance host resistance to infections and cancer.

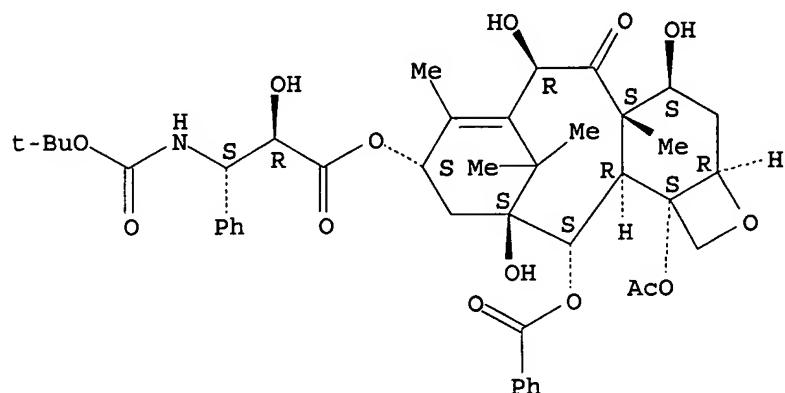
IT 114977-28-5, Taxotere

RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses) (combination with; free or liposome-encapsulated lipopeptide immunomodulators for tumor treatment and reduction of antitumor adverse effects)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 7 THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 35 OF 55 MEDLINE on STN
 ACCESSION NUMBER: 97356889 MEDLINE
 DOCUMENT NUMBER: PubMed ID: 9213325
 TITLE: Docetaxel combined with vinorelbine: phase I results and new study designs.
 AUTHOR: Fumoleau P; Fety R; Delecroix V; Perrocheau G; Azli N
 CORPORATE SOURCE: Medical Oncology Department, Centre Rene Gauducheau, CRLCC Nantes-Atlantique, Nantes-St Herblain, France.
 SOURCE: Oncology (Williston Park, N.Y.), (1997 Jun) Vol. 11, No. 6 Suppl 6, pp. 29-31.
 Journal code: 8712059. ISSN: 0890-9091.
 PUB. COUNTRY: United States
 DOCUMENT TYPE: (CLINICAL TRIAL)
 (CLINICAL TRIAL, PHASE I)
 Journal; Article; (JOURNAL ARTICLE)
 LANGUAGE: English
 FILE SEGMENT: Priority Journals
 ENTRY MONTH: 199709
 ENTRY DATE: Entered STN: 16 Sep 1997
 Last Updated on STN: 16 Sep 1997
 Entered Medline: 2 Sep 1997

AB This was a phase I dose-finding and pharmacokinetic study of vinorelbine (Navelbine) and docetaxel (Taxotere) as first-line chemotherapy for metastatic breast cancer. Vinorelbine dose, 20 or 22.5 mg/m², on days 1 and 5, was followed on day 1 by docetaxel every 21 days, in doses increasing from 60 to 100 mg/m². Two maximum tolerated doses were reached, the first at 75 mg/m² of docetaxel and 22.5 mg/m² of vinorelbine, and the second at 100 mg/m² of docetaxel and 20 mg/m² of vinorelbine. Symptomatic peripheral neuropathy was not observed.

The recommended doses for phase II studies are 75 to 85 mg/m² of docetaxel on day 1 and 20 mg/m² of vinorelbine on days 1 and 5, every 3 weeks. The treatment regimen, which included 3-day corticosteroid prophylaxis, resulted in only mild fluid retention. Responses were seen at all dose levels, with an 80% overall response rate at the higher recommended dose; the overall response rate for patients at all dose levels was 66%. A high rate of response, including a complete response, was observed in patients with liver metastases.

L12 ANSWER 36 OF 55 MEDLINE on STN
ACCESSION NUMBER: 96150144 MEDLINE
DOCUMENT NUMBER: PubMed ID: 8546908
TITLE: A late phase II study of RP56976 (docetaxel) in patients with advanced or recurrent breast cancer.
AUTHOR: Adachi I; Watanabe T; Takashima S; Narabayashi M; Horikoshi N; Aoyama H; Taguchi T
CORPORATE SOURCE: Department of Medical Oncology, National Cancer Center Hospital, Tokyo, Japan.
SOURCE: British journal of cancer, (1996 Jan) Vol. 73, No. 2, pp. 210-6.
PUB. COUNTRY: SCOTLAND: United Kingdom
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199602
ENTRY DATE: Entered STN: 6 Mar 1996
Last Updated on STN: 6 Feb 1998
Entered Medline: 16 Feb 1996
AB A late phase II clinical trial of RP56976 (docetaxel), derived from Taxus baccata was performed to evaluate anti-tumour activity, time to progression and clinical toxicity in patients with advanced or recurrent breast cancer. The patients, between 15 and 80 years old with performance status (PS) of 0-2, received at least two cycles of docetaxel 60 mg m⁻² intravenously at 3-4 week intervals. Of the 81 patients enrolled, the 72 eligible for the study were given a total of 327 cycles, with a median of four cycles each. Five patients obtained a complete response (CR) and 27 a partial response (PR); the response rate (RR) was 44.4% (95% confidence interval 32.7-56.6%). A relatively high RR of 9/28 (32.1%) was observed in patients who had received prior chemotherapy involving anthracyclines. The dose-limiting toxicity was grade 3-4 leucocytopenia or neutropenia, found in 78.9% and 85.9% patients respectively. Other severe (grade > 3) toxicities included alopecia (38%), anorexia (18.3%), nausea/vomiting (11.3%), and fatigue (9.9%). Hypersensitivity reactions, oedema and skin toxicity were not severe and were reversible. One therapy-related death occurred 10 days after the initial dose was given. These findings indicate that docetaxel has potent activity against metastatic breast cancer, and that the dose of 60 mg m⁻² is safe.

L12 ANSWER 37 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2002:123598 CAPLUS.
DOCUMENT NUMBER: 136:161350
TITLE: Method of inhibiting angiogenesis associated with malignant and neoplastic cells using active vitamin D analogs
INVENTOR(S): Bishop, Charles W.; Mazess, Richard B.
PATENT ASSIGNEE(S): Bone Care International, Inc., USA
SOURCE: U.S. Pat. Appl. Publ., 13 pp., Cont.-in-part of U.S. Ser. No. 596,149.
CODEN: USXXCO
DOCUMENT TYPE: Patent
LANGUAGE: English

FAMILY ACC. NUM. COUNT: 20

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-----------------|--------------|
| US 2002019375 | A1 | 20020214 | US 2001-891805 | 20010626 |
| US 6573256 | B2 | 20030603 | | |
| US 5763429 | A | 19980609 | US 1996-781910 | 19961230 <-- |
| US 6537982 | B1 | 20030325 | US 1998-596149 | 19980223 |
| PRIORITY APPLN. INFO.: | | | US 1996-781910 | A3 19961230 |
| | | | US 1998-596149 | A2 19980223 |
| | | | US 1993-119895 | A2 19930910 |
| | | | US 1994-265438 | A2 19940624 |
| | | | US 1995-415488 | A2 19950403 |
| | | | US 1995-486387 | A2 19950607 |

OTHER SOURCE(S): MARPAT 136:161350

AB Methods are disclosed which use active vitamin D analogs for the inhibition of angiogenesis associated with malignant and neoplastic cells. Methods comprise the application of an effective amount of a hypocalcemic hydroxyvitamin D compound to inhibit the angiogenesis of malignant cells, induce the apoptosis of malignant cells, and regress the growth of tumor cells.

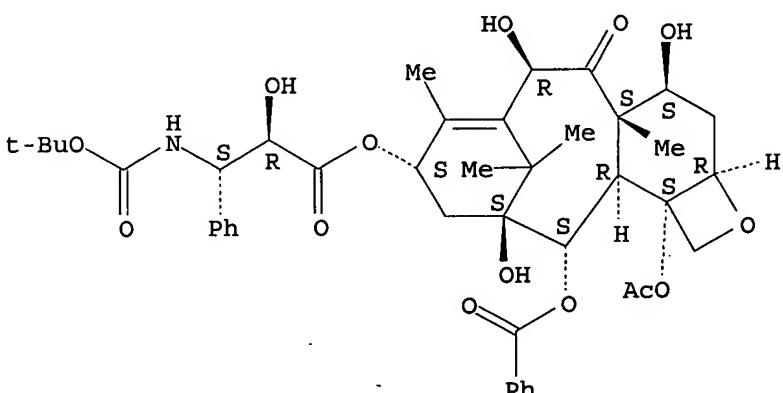
IT 114977-28-5, Docetaxel

RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(hydroxyvitamin D compds. for inhibition of tumor-associated angiogenesis, and use with other agents)

RN 114977-28-5 CAPLUS

CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 38 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2002:72803 CAPLUS

DOCUMENT NUMBER: 136:113175

TITLE: Method of treating malignancy-associated hypercalcemia using active vitamin D analogs

INVENTOR(S): Bishop, Charles W.; Mazess, Richard B.

PATENT ASSIGNEE(S): Bone Care International, Inc., USA

SOURCE: U.S. Pat. Appl. Publ., 12 pp., Cont.-in-part of U.S. 5,763,429.

CODEN: USXXCO

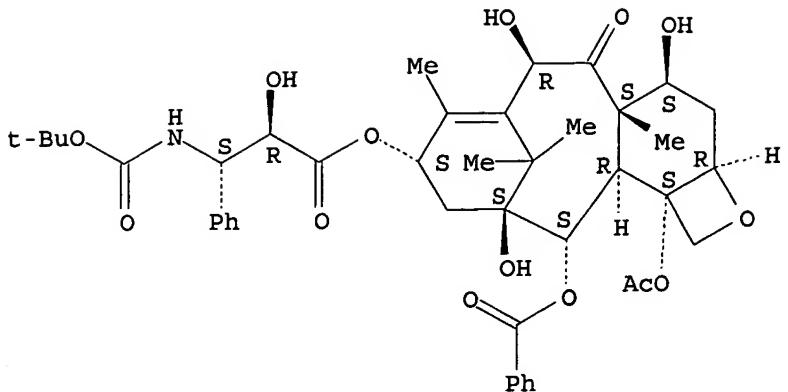
DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 20
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|-----------------|-----------------|--------------|
| US 2002010165 | A1 | 20020124 | US 2001-891763 | 20010626 |
| US 6566353 | B2 | 20030520 | | |
| US 5763429 | A | 19980609 | US 1996-781910 | 19961230 <-- |
| CA 2451037 | AA | 20030109 | CA 2002-2451037 | 20020626 |
| WO 2003002060 | A2 | 20030109 | WO 2002-US20320 | 20020626 |
| WO 2003002060 | A3 | 20040311 | | |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, UZ, VN, YU, ZA, ZM, ZW | | | | |
| RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG | | | | |
| EP 1416939 | A2 | 20040512 | EP 2002-747979 | 20020626 |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL, TR | | | | |
| CN 1520301 | A | 20040811 | CN 2002-812871 | 20020626 |
| JP 2004535439 | T2 | 20041125 | JP 2003-508302 | 20020626 |
| US 2003207810 | A1 | 20031106 | US 2003-441731 | 20030520 |
| PRIORITY APPLN. INFO.: | | | | |
| | | US 1996-781910 | A2 19961230 | |
| | | US 1993-119895 | A2 19930910 | |
| | | US 1994-265438 | A2 19940624 | |
| | | US 1995-415488 | A2 19950403 | |
| | | US 1995-486387 | A2 19950607 | |
| | | US 1998-596149 | A3 19980223 | |
| | | US 2001-891763 | A 20010626 | |
| | | WO 2002-US20320 | W 20020626 | |

OTHER SOURCE(S): MARPAT 136:113175

- AB Methods utilizing active vitamin D analogs for the treatment of malignancy-associated hypercalcemia. Methods comprise the application of an effective amount of a hypocalcemic vitamin D compound to alleviate hypercalcemia, lower serum parathyroid hormone related protein (PTHrP) levels. The hypocalcemic vitamin D compds. can be coadministered with a cytotoxic agent.
- IT 114977-28-5, Docetaxel
 RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (method of treating malignancy-associated hypercalcemia using active vitamin D analogs coadministered with cytotoxic agents)
- RN 114977-28-5 CAPLUS
- CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9;10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



L12 ANSWER 39 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN

ACCESSION NUMBER: 2001:507531 CAPLUS

DOCUMENT NUMBER: 135:107247

TITLE: Preparation of 3-heteroarylidenyl-2-indolinone compounds for modulating protein kinase activity and for use in cancer chemotherapy

INVENTOR(S): Langecker, Peter J.; Shawver, Laura K.; Tang, Peng C.; Sun, Li

PATENT ASSIGNEE(S): Sugen, Inc., USA

SOURCE: PCT Int. Appl., 93 pp.

CODEN: PIXXD2

DOCUMENT TYPE: Patent

LANGUAGE: English

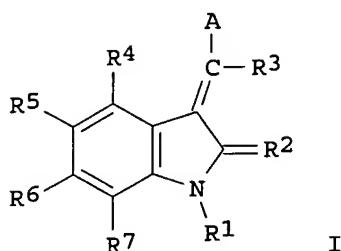
FAMILY ACC. NUM. COUNT: 2

PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------|--------------|
| WO 2001049287 | A1 | 20010712 | WO 2000-US18058 | 20000630 |
| W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CR, CU, CZ, DE, DK, DM, DZ, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZW
RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| WO 2000038519 | A1 | 20000706 | WO 1999-US31232 | 19991230 <-- |
| W: AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, CA, CH, CN, CU, CZ, DE, DK, EE, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MD, MG, MK, MN, MW, MX, NO, NZ, PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TR, TT, UA, UG, UZ, VN, YU, ZW
RW: GH, GM, KE, LS, MW, SD, SL, SZ, TZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| US 2003073837 | A1 | 20030417 | US 1999-476232 | 19991230 |
| EP 1259234 | A1 | 20021127 | EP 2000-943334 | 20000630 |
| EP 1259234 | B1 | 20060816 | | |
| R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT, IE, SI, LT, LV, FI, RO, MK, CY, AL | | | | |
| JP 2003535038 | T2 | 20031125 | JP 2001-549655 | 20000630 |
| US 2003191162 | A1 | 20031009 | US 2002-307483 | 20021202 |
| PRIORITY APPLN. INFO.: | | | US 1999-476232 | A 19991230 |
| | | | WO 1999-US31232 | A 19991230 |
| | | | US 2000-569545 | A 20000512 |
| | | | US 1998-114313P | P 19981231 |

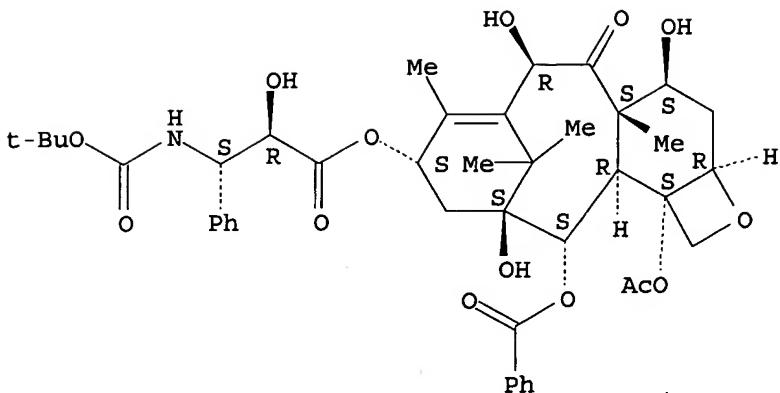
OTHER SOURCE(S) :
GI

MARPAT 135:107247



- AB** The present invention relates to 3-heteroarylidenyl-2-indolinone compds. [I; R1 = H, alkyl; R2 = O, S; R3 = H; R4 , R5, R6, R7 = H, alkyl, alkoxy, aryl, aryloxy, alkaryloxy, halo, trihalomethyl, S(O)R, SO2NRR', SO3R, SR, NO2, NRR', OH, cyano, COR, O2CR, (CH2)nCO2R, CONRR'; A = a five membered heteroaryl selected from (un)substituted thiophene, pyrrole, pyrazole, imidazole, 1,2,3-triazole, 1,2,4-triazole, oxazole, isoxazole, thiazole, isothiazole, 2-sulfonylfuran, 4-alkylfuran, 1,2,3-oxadiazole, 1,2,4-oxadiazole, 1,2,5-oxadiazole, 1,3,4-oxadiazole, 1,2,3,4-oxatriazole, 1,2,3,5-oxatriazole, 1,2,3-thiadiazole, etc.; n = 0-3; R, R' = H, alkyl, aryl] or physiol. acceptable salts or prodrugs thereof are prepared. These compds. modulate the enzymic activity of protein kinases such as receptor protein tyrosine kinase, cellular tyrosine kinase, and serine threonine kinase and therefore are expected to be useful in the prevention and treatment of protein kinase related cellular disorders such as cancer. Furthermore, these compds. are expected to enhance the efficacy of other chemotherapeutic agents, in particular, fluorinated pyrimidines, in the treatment of cancer. In a cellular-based assay for inhibiting the receptor phosphorylation, 3-[(2,4-dimethylpyrrol-5-yl)methylidenyl]-2-indolinone (II) inhibited Flk-1-autophosphorylation with IC50 of .apprx.1 μM. II in vitro inhibited proliferation of endothelial cells induced by VEGF with IC50 of .apprx.0.07 μM. Although II in vitro had no direct inhibitory effect on a variety of tumor cell lines at concentration up to 50 μM, it in vivo demonstrated a significant suppression of tumor growth against a broad spectrum of tumor types s.c. implanted into immunocompromised mice and whose growth are driven by various growth factors such as PDGF, EGF, and Her2.
- IT** 114977-28-5, Docetaxel
RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
(for cancer chemotherapy in combination with heteroarylidenylindolinone derivative; preparation of 3-heteroarylidenyl-2-indolinone compds. for modulating protein kinase activity for cancer chemotherapy)
- RN** 114977-28-5 CAPLUS
CN Benzenepropanoic acid, β-[[[(1,1-dimethylethoxy)carbonyl]amino]-α-hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetyloxy)-12-(benzyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (αR,βS)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 8 THERE ARE 8 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 40 OF 55 MEDLINE on STN

ACCESSION NUMBER: 2000024226 MEDLINE

DOCUMENT NUMBER: PubMed ID: 10560436

TITLE: A case of hepatic arterial infusion chemotherapy with docetaxel for liver metastasis from breast cancer.

AUTHOR: Kim S J; Maeura Y; Ueda N; Saito M; Matsunaga S

CORPORATE SOURCE: Senri Hoken Medical Center, Dept. of Surgery, Shinsenri Hospital.

SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1999 Oct) Vol. 26, No. 12, pp. 1959-62.
Journal code: 7810034. ISSN: 0385-0684.

PUB. COUNTRY: Japan

DOCUMENT TYPE: (CASE REPORTS)

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Japanese

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199911

ENTRY DATE: Entered STN: 13 Jan 2000

Last Updated on STN: 13 Jan 2000

Entered Medline: 26 Nov 1999

AB We experienced a case of hepatic arterial infusion chemotherapy using docetaxel for liver metastasis, which showed no response to CEF therapy, from breast cancer. A 63-year-old woman had undergone modified radical mastectomy for right breast cancer (T2aN1bM0: Stage II) in October, 1995. Six-cycle CMF therapy and toremifene citrate (40 mg/day)

were administered as adjuvant therapy, but multiple recurrent tumors in liver, lung, and local site were detected in February 1997. Six-cycle CEF therapy was given for recurrent disease and there was a complete response for lung and local recurrence, but no change in liver metastasis.

Chemoendocrine therapies using 5'-DFUR or CMitF in addition to TAM and fadrozole hydrochloride hydrate had developed progressive disease for liver metastasis. A catheter and port kit were operatively inserted and implanted in March 1998. Hepatic arterial infusion of docetaxel (30-40 mg/body/month, one hour administration) was repeated 4 times, once in our clinic. Leukopenia, general fatigue and fever, which were mild and did not require any treatment, appeared as side effects. This treatment reduced multiple liver metastatic sites on abdominal CT finding and was thought to be a partial response. However, the patient had multiple brain metastasis and died on August 2, 1998. While docetaxel, even by systemic administration, has a 36-77% response rate for liver metastasis, arterial infusion might have a good response and mild side effect with a lower dose than by intravenous administration.

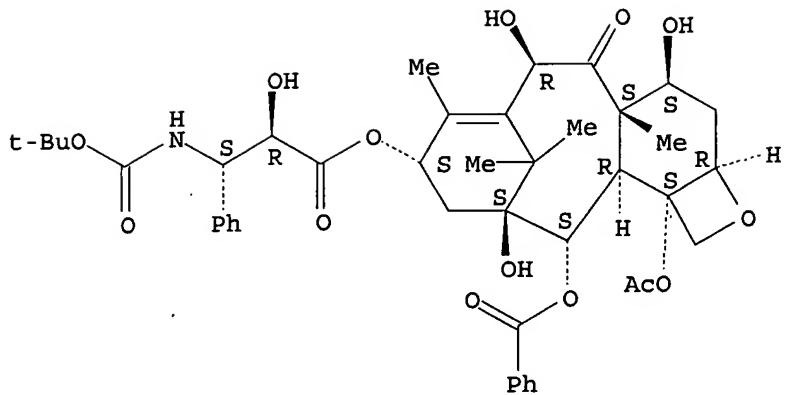
L12 ANSWER 41 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 2001:161407 CAPLUS

DOCUMENT NUMBER: 134:202681
 TITLE: Dietary supplementation with, and methods for,
 administration of a yeast-derived selenium product,
 and use in cancer chemotherapy
 INVENTOR(S): Hsia, Houn Simon; Yang, Ping; Arnold, Michael
 PATENT ASSIGNEE(S): Viva America Marketing Corporation, USA
 SOURCE: U.S., 9 pp., Cont.-in-part of U.S. 6,140,107.
 CODEN: USXXAM
 DOCUMENT TYPE: Patent
 LANGUAGE: English
 FAMILY ACC. NUM. COUNT: 4
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|------------------------|------|----------|-------------------------|--------------|
| US 6197295 | B1 | 20010306 | US 1999-303993 | 19990503 |
| US 6140107 | A | 20001031 | US 1996-719572 | 19960925 <-- |
| US 6368643 | B1 | 20020409 | US 1999-298114 | 19990423 |
| US 2001043925 | A1 | 20011122 | US 2001-801124 | 20010305 |
| US 6576233 | B2 | 20030610 | | |
| PRIORITY APPLN. INFO.: | | | | |
| | | | US 1996-719572. A2 | 19960925 |
| | | | US 1997-802773 | B2 19970221 |
| | | | US 1998-15758 | A2 19980129 |
| | | | US 1998-82939P | P 19980424 |
| | | | US 1999-303993 | A3 19990503 |

- AB The invention solves the need for nontoxic forms of selenium which is an essential part of the human diet. The invention provides dried-yeast products containing selenium, as well as a method of producing the dried yeast products. The method uses selenium having high biol. activity but low toxicity. The invention also provides nutritional supplements containing the selenium-containing dried yeast products and methods of administering these products and supplements to improve human health. The invention also provides a practically nontoxic yeast selenium product having increased intracellular selenium concns., as well as methods to reduce tumor cell growth by administration of a selenium yeast product comprising yeast *Saccharomyces boulardii* sequela PY31 (ATCC 74366) in combination with chemotherapeutic agents.
- IT 114977-28-5, Taxotere
 RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
 (dietary supplementation with yeast-derived selenium product, and use in cancer chemotherapy)
- RN 114977-28-5 CAPLUS
- CN Benzenepropanoic acid, β -[[[(1,1-dimethylethoxy)carbonyl]amino]- α -hydroxy-, (2aR,4S,4aS,6R,9S,11S,12S,12aR,12bS)-12b-(acetoxy)-12-(benzoyloxy)-2a,3,4,4a,5,6,9,10,11,12,12a,12b-dodecahydro-4,6,11-trihydroxy-4a,8,13,13-tetramethyl-5-oxo-7,11-methano-1H-cyclodeca[3,4]benz[1,2-b]oxet-9-yl ester, (α R, β S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 42 OF 55 MEDLINE on STN
 ACCESSION NUMBER: 2001191709 MEDLINE
 DOCUMENT NUMBER: PubMed ID: 11124653
 TITLE: Chemotherapy-induced noncardiogenic pulmonary edema related to gemcitabine plus docetaxel combination with granulocyte colony-stimulating factor support.
 AUTHOR: Briassoulis E; Froudarakis M; Milionis H J; Peponis I; Constantopoulos S; Pavlidis N
 CORPORATE SOURCE: Department of Medical Oncology, Ioannina University Hospital, Ioannina, Greece.. ebriasou@otenet.gr
 SOURCE: Respiration; international review of thoracic diseases, (2000) Vol. 67, No. 6, pp. 680-3.
 Journal code: 0137356. ISSN: 0025-7931.
 PUB. COUNTRY: Switzerland
 DOCUMENT TYPE: (CASE REPORTS)
 Journal; Article; (JOURNAL ARTICLE)
 LANGUAGE: English
 FILE SEGMENT: Priority Journals
 ENTRY MONTH: 200104
 ENTRY DATE: Entered STN: 10 Apr 2001
 Last Updated on STN: 10 Apr 2001
 Entered Medline: 5 Apr 2001

AB Several cancer therapeutic agents have been associated with pulmonary toxicity. Herein, we describe the case of a 73-year-old woman with breast cancer metastatic to the liver, who developed noncardiogenic pulmonary edema (NPE) while on treatment with gemcitabine plus docetaxel combination with granulocyte colony-stimulating factor (G-CSF) support. Gemcitabine, a deoxycytidine analogue, is reported to produce mild self-limiting and only occasionally severe pulmonary toxicity. The microtubule stabilizer docetaxel has been associated with water retention complications. The combination of these two agents has shown promising activity in several solid tumors and is in a phase of clinical development with prophylactic G-CSF in most of the trials due to the high rate of dose-limiting neutropenia observed with this combination. In our case pulmonary toxicity resolved rapidly following the administration of corticosteroids. A possible deleterious synergy of the compounds involved in this case is discussed and the medical literature on NPE related to cancer therapy is shortly reviewed. We conclude that NPE should always be considered in patients with respiratory function deterioration while on therapy with the gemcitabine-docetaxel combination and G-CSF. Corticosteroids can provide maximum benefit if started early upon diagnosis coupled with withdrawal of the causative drugs.

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ACCESSION NUMBER: 1999197818 MEDLINE
DOCUMENT NUMBER: PubMed ID: 10097745
TITLE: A late phase II clinical study of RP56976 (docetaxel) in patients with advanced or recurrent gastric cancer: a cooperative study group trial (group B).
AUTHOR: Mai M; Sakata Y; Kanamaru R; Kurihara M; Suminaga M; Ota J; Hirabayashi N; Taguchi T; Furue H
CORPORATE SOURCE: Dept. of Surgery, Cancer Research Institute, Kanazawa University.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1999 Mar) Vol. 26, No. 4, pp. 487-96.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
(MULTICENTER STUDY)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199904
ENTRY DATE: Entered STN: 13 Apr 1999..
Last Updated on STN: 13 Apr 1999
Entered Medline: 1 Apr 1999

AB A late phase II clinical study of RP56976 (docetaxel) in patients with advanced or recurrent gastric cancer was performed to evaluate the anti-tumor activity and clinical toxicity as a multicenter cooperative trial. Docetaxel was administered intravenously at a dose of 60 mg/m² every 3-4 weeks. Of 72 patients enrolled, 63 patients were eligible and 59 patients were evaluable for response. The anti-tumor effects obtained complete response (CR) in one patient partial response (PR) in 13, minor response (MR) in 3, no change (NC) in 20, and progressing disease (PD) in 22 patients. The overall response rate in 59 patients was 23.7% (14/59). For 14 CR or PR cases, a response appeared 10 to 107 days (median 33.5 days) and 1 to 8 (median 2) times of dosing after the initial administration. The response rate was 9.5% in the primary tumor, 31.3% livers, 50.0% abdominal tumor, and 24.1% lymph nodes, respectively. The major adverse reactions were gastrointestinal symptoms including nausea/vomiting, anorexia, fatigue, alopecia and fever. Leukocytopenia and neutrocytopenia were also observed with a high incidence, but they recovered after 8 days from the nadir. The results show that docetaxel is an effective anti-tumor agent for advanced or recurrent gastric cancer. It is necessary to conduct another clinical trial by concomitant administration with other anti-tumor agents.

L12 ANSWER 44 OF 55 MEDLINE on STN
ACCESSION NUMBER: 1999014548 MEDLINE
DOCUMENT NUMBER: PubMed ID: 9797814
TITLE: Late phase II clinical study of RP56976 (docetaxel) in patients with advanced/recurrent gastric cancer: a Japanese Cooperative Study Group trial (group A).
AUTHOR: Taguchi T; Sakata Y; Kanamaru R; Kurihara M; Suminaga M; Ota J; Hirabayashi N
CORPORATE SOURCE: Japan Society for Cancer Chemotherapy, Aomori Prefectural Central Hospital.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1998 Oct) Vol. 25, No. 12, pp. 1915-24.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
(MULTICENTER STUDY)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals

ENTRY MONTH: 199811
ENTRY DATE: Entered STN: 6 Jan 1999
Last Updated on STN: 6 Jan 1999
Entered Medline: 4 Nov 1998

AB A late phase II clinical study of RP56976 (docetaxel) was conducted in patients with advanced/recurrent gastric cancer as a multicenter cooperative trial. Docetaxel was administered intravenously at a dose of 60 mg/m² every 3-4 weeks. Of the 76 patients enrolled, 66 patients were eligible and 59 patients were evaluable for response. One patient showed complete response (CR), 13 patients partial response (PR), 1 patient minor response (MR), 19 patients no change (NC) and 25 patients had progressive disease (PD). The overall response rate in 59 evaluable patients was 23.7% (95% CI = 13.6-36.6%). The primary tumor showed a 4.3% (1/23) response, while the metastatic lesions in the abdomen, pelvic mass, lung, liver, and lymph nodes showed response rates of 62.5% (5/8), 33.3% (1/3), 33.3% (1/3), 14.8% (4/27), and 13.9% (5/26), respectively. About hematological toxicity, severe (Grade 3 or more) leukopenia was observed in 36 patients (56.3%) and neutropenia in 52 patients (81.3%). Other major toxicity (Grade 3 or more) included nausea/vomiting in 11 patients (17.2%), anorexia in 9 patients (14.1%), fatigue in 5 patients (7.8%), and alopecia in 7 patients (10.9%), all which were tolerable. The results show that docetaxel is an effective anticancer agent for advanced/recurrent gastric cancer.

L12 ANSWER 45 OF 55 MEDLINE on STN
ACCESSION NUMBER: 2000464139 MEDLINE
DOCUMENT NUMBER: PubMed ID: 11016005
TITLE: A case of effective chemotherapy using CAF followed by docetaxel for advanced breast cancer.
AUTHOR: Kokufu I; Taniguchi H; Kim Y H; Fukuda K; Yamamoto M; Yano T; Yamada K; Kitano H; Fukuda H
CORPORATE SOURCE: Dept. of Surgery, Itami City Hospital.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (2000 Sep) Vol. 27, No. 10, pp. 1577-80.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CASE REPORTS)
Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200010
ENTRY DATE: Entered STN: 19 Oct 2000
Last Updated on STN: 19 Oct 2000
Entered Medline: 10 Oct 2000

AB A huge mass measuring 13 x 12 cm and wide cutaneous edema were detected in the right breast of a 51-year-old woman. Under a diagnosis of locally advanced breast cancer (T4bN2M1, stage IV) with liver metastases, we attempted sequential neoadjuvant chemotherapy. After three courses of CAF therapy (cyclophosphamide, doxorubicin (DXR), 5-FU), the primary tumor was decreased by 56% and the liver metastases had disappeared. A minor pathologic response was observed. Subsequently, three courses of docetaxel (TXT) administration were carried out. The primary tumor was then decreased by 75% and the axillary metastases had disappeared. Histopathological examination showed gross viable tumor cells in the residual tumor and positive axillary lymph nodes. The only toxic effect was nausea (grade 1) and no major adverse effects were observed. Neoadjuvant chemotherapy with sequential DXR followed by TXT is a useful treatment for locally advanced breast cancer.

L12 ANSWER 46 OF 55 MEDLINE on STN
ACCESSION NUMBER: 1999430325 MEDLINE
DOCUMENT NUMBER: PubMed ID: 10500538
TITLE: A case of recurrent breast cancer successfully treated with docetaxel.

AUTHOR: Koshizuka K; Hada M; Muto S; Hagiwara J; Nakagomi H; Takano K; Kamiya K; Tada Y
CORPORATE SOURCE: Second Dept. of Surgery, Yamanashi Medical University.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1999 Sep) Vol. 26, No. 10, pp. 1479-81.
Journal code: 7810034. ISSN: 0385-0684.

PUB. COUNTRY: Japan
DOCUMENT TYPE: (CASE REPORTS)
Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: Japanese
FILE SEGMENT: Priority Journals

ENTRY MONTH: 199909
ENTRY DATE: Entered STN: 12 Oct 1999
Last Updated on STN: 12 Oct 1999
Entered Medline: 30 Sep 1999

AB A 53-year-old female underwent mastectomy for left breast cancer in April, 1993. She was given oral tamoxifen but this had to be discontinued due to its side effects. In March, 1998, she developed bone and lung metastases, in spite of treatment with combination chemotherapy (CEF). We thus treated here with docetaxel 90 mg three times and 40 mg six times. After the chemotherapy, she achieved complete remissions of the lung metastases and a decrease in serum CEA, CA 15-3, NCC-ST439, and BCA225. Adverse reactions to docetaxel were grade 2 alopecia, grade 4 neutropenia, dysgeusia, and fluid retention. All were tolerable. This new agent may play an important future role in chemotherapy for recurrent breast cancer.

L12 ANSWER 47 OF 55 MEDLINE on STN
ACCESSION NUMBER: 1998233482 MEDLINE
DOCUMENT NUMBER: PubMed ID: 9571974
TITLE: Breast cancer with liver metastasis responsive to docetaxel: case report.
AUTHOR: Oura S; Sakurai T; Yoshimura G; Tamaki T; Umemura T; Kokawa Y
CORPORATE SOURCE: Dept. of Surgery, Wakayama Medical College Kihoku Hospital.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (1998 Apr) Vol. 25, No. 5, pp. 743-6.
Journal code: 7810034. ISSN: 0385-0684.

PUB. COUNTRY: Japan
DOCUMENT TYPE: (CASE REPORTS)
Journal; Article; (JOURNAL ARTICLE)
LANGUAGE: Japanese
FILE SEGMENT: Priority Journals
ENTRY MONTH: 199805
ENTRY DATE: Entered STN: 14 May 1998
Last Updated on STN: 14 May 1998
Entered Medline: 7 May 1998

AB A 59-year-old female underwent mastectomy for right breast cancer in November 1992. She received tamoxifen and anthracycline-containing chemotherapy as adjuvant therapy. In and after September 1994, she developed loco-regional recurrences five times in total, each of which was treated with surgery and conventional combination chemotherapy. In April 1997, she developed liver metastasis, which was refractory to biochemical modulation therapy (low-dose cisplatin + 5-FU). We, therefore, treated her six times with docetaxel 80 mg, which resulted in partial response of the liver metastasis and brought about a marked decrease in serum CA15-3 levels. Adverse effects of docetaxel were grade 3 alopecia and leucocytopenia. She has been well without re-growth of the liver metastasis for over five months.

L12 ANSWER 48 OF 55 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
ACCESSION NUMBER: 1998:259786 BIOSIS
DOCUMENT NUMBER: PREV199800259786

TITLE: Antitumour effect of docetaxel in malignant diseases.
 AUTHOR(S): Eckhardt, Sandor [Reprint author]
 CORPORATE SOURCE: Rath Gyorgy u. 7-9, 1122 Budapest, Hungary
 SOURCE: Orvosi Hetilap, (April 12, 1998) Vol. 139, No. 15, pp. 867-872. print.
 CODEN: ORHEAG. ISSN: 0030-6002.
 DOCUMENT TYPE: Article
 General Review; (Literature Review)
 LANGUAGE: Hungarian
 ENTRY DATE: Entered STN: 9 Jun 1998
 Last Updated on STN: 12 Aug 1998
 AB In recent years numerous molecular biological discoveries enlightened the various steps of the neoplastic transformation. Based on new targets, this development made it possible to synthesize new tumour inhibitory substances. Among them taxanes capable to block depolymerization of tubulin - which is an essential molecule in cell division - play an important role. Docetaxel (Taxotere) belongs to this group and is an active drug in the treatment of breast cancer. Moreover, platinum-resistant tumours may also respond to the therapy. It is important to note that even visceral (hepatic) metastases may express chemosensitivity. Results of combination chemotherapy seem to be also promising. The antitumour effect of Taxotere in NSCLC and other malignant neoplasms is under investigation. The toxicity of Taxotere may be successfully reduced by premedication of steroids. The necessary protective measures render the Taxotere therapy safe and of being perspectivistic.

L12 ANSWER 49 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
 ACCESSION NUMBER: 1999:325975 CAPLUS
 DOCUMENT NUMBER: 130:357177
 TITLE: Detoxication of active pharmaceutical substances using cyclodextrin oligomers
 INVENTOR(S): Moser, Joerg G.
 PATENT ASSIGNEE(S): Germany
 SOURCE: PCT Int. Appl., 31 pp.
 CODEN: PIXXD2
 DOCUMENT TYPE: Patent
 LANGUAGE: German
 FAMILY ACC. NUM. COUNT: 1
 PATENT INFORMATION:

| PATENT NO. | KIND | DATE | APPLICATION NO. | DATE |
|---|------|----------|-----------------------------|--------------|
| WO 9924474 | A1 | 19990520 | WO 1998-EP7229 | 19981111 <-- |
| W: AL, AU, BA, BB, BG, BR, CA, CN, CU, CZ, EE, GE, HR, HU, ID, IL, IS, JP, KP, KR, LC, LK, LR, LS, LT, LV, MG, MK, MN, MX, NO, NZ, PL, RO, SG, SI, SK, SL, TR, TT, UA, US, UZ, VN, YU, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM | | | | |
| RW: GH, GM, KE, LS, MW, SD, SZ, UG, ZW, AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, BF, BJ, CF, CG, CI, CM, GA, GN, GW, ML, MR, NE, SN, TD, TG | | | | |
| AU 9916694 | A1 | 19990531 | AU 1999-16694 | 19981111 <-- |
| EP 1045863 | A1 | 20001025 | EP 1998-961184 | 19981111 <-- |
| EP 1045863 | B1 | 20030402 | | |
| R: AT, BE, CH, DE, ES, FR, GB, IT, LI, NL | | | | |
| JP 2001522901 | T2 | 20011120 | JP 2000-520482 | 19981111 |
| AT 236195 | E | 20030415 | AT 1998-961184 | 19981111 |
| US 6642214 | B1 | 20031104 | US 2000-554223 | 20000803 |
| PRIORITY APPLN. INFO.: | | | DE 1997-19749801 A 19971111 | |
| | | | DE 1998-19822416 A 19980519 | |
| | | | WO 1998-EP7229 W 19981111 | |

AB Cyclodextrin oligomers with 2 cyclodextrins connected via a spacer B on the secondary side [CD-X-A-X-B-X-A-X-CD; CD = cyclodextrin; X = bond, NH,

O, S, C(O); A = bond, C2-4 aliphatic residue; B = rigid, preferably hydrophilic residue] form strongly hydrophilic inclusion compds. with pharmaceutical agents and thereby prevent toxic side effects of drugs on nontarget cells by inhibiting their uptake into the cells. The drugs can be targeted to specific tissue sites by attachment of affinity groups such as antibodies to the cyclodextrin residues, and the drug can be released at the target site by destruction of the cyclodextrin residues (e.g. with cyclodextrinase from Klebsiella oxytoca). Provided the cyclodextrins are connected on their secondary sides, their cavities will face each other; the distance between them is determined by the choice of spacer, and is preferably 0.8-1.8 nm. Thus, β -cyclodextrin was condensed with 4,4'-methylenebis(benzenesulfonyl chloride) and the product reacted with diaminopropane to form β -6(A-D)-diamidopropanediaminocyclodextrin (I). Sep., 2-monotosyl- β -cyclodextrin reacted with 3-mercaptopropionic acid to form β -(2)cyclodextrin-(3-thiopropionic acid) (II). Reaction of II with carbonyldimidazole, N-hydroxysuccinimide, and a 2.5-fold molar excess of I produced a cyclodextrin trimer. Nude mice bearing OAT SCLC cell tumors were treated with biotinylated monoclonal antibody ICO 25 i.p., followed 24 h later by NeutrAvidin i.p., and after an addnl. 48 h by a biotinylcadaverine-labeled CD dimer-paclitaxel complex.. Growth of the tumors was inhibited without occurrence of side effects.

REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 50 OF 55 CAPLUS COPYRIGHT 2006 ACS on STN
ACCESSION NUMBER: 1998:213711 CAPLUS
DOCUMENT NUMBER: 128:289570
TITLE: Pharmacokinetics of anticancer agents in patients with impaired liver function
AUTHOR(S): Donelli, M. G.; Zucchetti, M.; Munzone, E.; D'incalci, M.; Crosignani, A.
CORPORATE SOURCE: Dipartimento di Oncologia, Istituto di Ricerche Farmacologiche Mario Negri, Milan, 20157, Italy
SOURCE: European Journal of Cancer (1998), 34(1), 33-46
CODEN: EJCAEL; ISSN: 0959-8049
PUBLISHER: Elsevier Science Ltd.
DOCUMENT TYPE: Journal; General Review
LANGUAGE: English

AB A review with 95 refs. This report reviews published information on the clin. pharmacokinetics of antitumor agents in patients with liver dysfunction, associated with primary liver disease or liver metastases. Information was available for anthracyclines and their related compds., antimetabolites, cyclophosphamide, vinca alkaloids, taxanes and epipodophyllotoxins. Changes in the pharmacokinetic profile or metabolism in patients with mild or severe hepatobiliary dysfunction are described and the relationships between serum levels, parameters employed for measuring hepatic function and toxic or therapeutic effects are examined. Current knowledge of the pharmacokinetics of antineoplastic agents in liver disease is far from complete, mostly obtained in small nos. of non-homogeneous patients often presenting only moderate liver dysfunction, and empirical guidelines for dose assessment are still largely applied in clin. practice. Because of the complex pathophysiol. mechanisms of liver insufficiency in cancer patients, there is still doubt whether endogenous markers are useful. Although caution in treating cancer patients with liver insufficiency is compulsory, for most compds. there seems no need to recommend dose redns. for moderate impairment. However, for the tubulin acting agents, vincristine, vinblastine and possibly for paclitaxel and docetaxel, there is strong evidence that dose adjustment is mandatory in order to avoid excessive neutropenia and neurotoxicity.

REFERENCE COUNT: 95 THERE ARE 95 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 51 OF 55 MEDLINE on STN
ACCESSION NUMBER: 2000390579 MEDLINE
DOCUMENT NUMBER: PubMed ID: 10895201
TITLE: Preliminary results of multicenter phase II trial of docetaxel (Taxotere) in combination with doxorubicin as first line chemotherapy in Indonesian patients with advanced or metastatic breast cancer.
AUTHOR: Muthalib A; Darwis I; Prayogo N; Sutjipto
CORPORATE SOURCE: Dharmais National Cancer Center/School of Medicine, University of Indonesia, Jakarta.
SOURCE: Gan to kagaku ryoho. Cancer & chemotherapy, (2000 May) Vol. 27 Suppl 2, pp. 498-504.
Journal code: 7810034. ISSN: 0385-0684.
PUB. COUNTRY: Japan
DOCUMENT TYPE: (CLINICAL TRIAL)
(CLINICAL TRIAL, PHASE II)
Journal; Article; (JOURNAL ARTICLE)
(MULTICENTER STUDY)
(RANDOMIZED CONTROLLED TRIAL)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200008
ENTRY DATE: Entered STN: 18 Aug 2000
Last Updated on STN: 18 Aug 2000
Entered Medline: 10 Aug 2000

AB RATIONALE: Docetaxel and doxorubicin have produced a high degree of activity in previously untreated/treated patients with metastatic breast cancer (MBC). The efficacy of Taxotere (T) single agent as 2nd line chemotherapy is well established in large randomized phase III studies. OBJECTIVE: The objective of this study is to confirm the efficacy and safety of a combination of Taxotere with doxorubicin as 1st line chemotherapy in Indonesian MBC patients. TREATMENT AND METHOD: Eighteen patients age < or = 70 years with advanced or metastatic breast cancer (MBC) with no prior taxane chemotherapy or prior cumulative doxorubicin (D) of no more than 250 mg/m² and no heart disease were enrolled in this phase II study of D (50 mg/m²) IV bolus followed one hour later by Taxotere (T) 60 mg/m² IV infusion over 1 hour every 3 weeks for 6 cycles treatments. A 3-day oral corticosteroid premedication was administered starting one day before the infusion of each cycle. Left ventricular ejection fraction (LVEF) was evaluated at baseline and after cycle 6. PATIENTS CHARACTERISTICS: 18 patients (pts) have been treated with 108 cycles administered. Median age was 46 years (31-58), WHO PS 0 = 50%, 1 = 50% and number of organs involved were: 2 (72%), 3 (22%) and 4 (6%). RESULTS: After 3 cycles, partial (PR) and no change (NC) responses occurred in 15 pts (83.3%) and 3 pts (16.7%). The best overall response after 6 cycles, including complete (CR) and partial (PR) responses, occurred in 13 pts (72.2%) including 3 CRs and 10 PRs. Two patients with extensive liver metastases at the baseline had a complete disappearance after 6 cycles. No patients developed congestive heart failure (CHF). Grade 3/4 hematological toxicities included leukopenia in 18 pts (100%), febrile neutropenia in 6 pts (33%), leukopenia with infection in 2 pts (11%), leukopenia with fever in 1 pt (5.5%), and anemia in 6 pts (33.3%). Nonhematological toxicities grade 3/4 included alopecia (61%), asthenia (4.6%), nausea/vomiting (2.7%), pain (2.7%), stomatitis (2.7%), and diarrhoea (0.9%). Leukopenia was generally of short duration, occurred mainly during the first and second cycle, and did not require any dose reduction. There was one death due to progressive disease after six cycles of treatment. CONCLUSION: Taxotere--doxorubicin combination is very active in the first-line treatment of MBC, seems to be especially effective in patients with liver metastases, and is associated with a manageable toxicity profile.

ACCESSION NUMBER: 2001:89699 BIOSIS
DOCUMENT NUMBER: PREV200100089699
TITLE: Phase I study of weekly docetaxel in combination with capecitabine in patients with solid malignancies.
AUTHOR(S): Villalona-Calero, M. A. [Reprint author]; Shapiro, C. [Reprint author]; Otterson, G. A. [Reprint author]; Hauger, M. [Reprint author]; Kraut, E. [Reprint author]; Clinton, S. [Reprint author]; Shah, M. [Reprint author]; Stanek, M. [Reprint author]; Monk, J. P. [Reprint author]
CORPORATE SOURCE: Arthur James Cancer Center and R Solove Research Institute, Ohio State University, Columbus, OH, USA
SOURCE: Breast Cancer Research and Treatment, (November, 2000) Vol. 64, No. 1, pp. 125. print.
Meeting Info.: 23rd Annual San Antonio Breast Cancer Symposium. San antonio, Texas, USA. December 06-09, 2000. Cancer Therapy and Research Center Research Foundation.
CODEN: BCTR6. ISSN: 0167-6806.
DOCUMENT TYPE: Conference; (Meeting)
Conference; Abstract; (Meeting Abstract)
LANGUAGE: English
ENTRY DATE: Entered STN: 14 Feb 2001
Last Updated on STN: 12 Feb 2002

L12 ANSWER 53 OF 55 MEDLINE on STN
ACCESSION NUMBER: 2000339901 MEDLINE
DOCUMENT NUMBER: PubMed ID: 10885392
TITLE: Metastasectomy as a cytoreductive strategy for treatment of isolated pulmonary and hepatic metastases from breast cancer.
AUTHOR: Bathe O F; Kaklamanos I G; Moffat F L; Boggs J; Franceschi D; Livingstone A S
CORPORATE SOURCE: Department of Surgery, University of Miami, FL 33136, USA.. bathe@worldnet.att.net
SOURCE: Surgical oncology, (1999 Jul) Vol. 8, No. 1, pp. 35-42. Ref: 45
Journal code: 9208188. ISSN: 0960-7404.
PUB. COUNTRY: Netherlands
DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)
General Review; (REVIEW)
LANGUAGE: English
FILE SEGMENT: Priority Journals
ENTRY MONTH: 200007
ENTRY DATE: Entered STN: 28 Jul 2000
Last Updated on STN: 28 Jul 2000
Entered Medline: 20 Jul 2000

AB The authors sought to examine the utility of resection in conjunction with adjuvant chemotherapy for treatment of metastases from breast cancer isolated to the liver or lungs. Limitations of regional therapy were examined and potential agents for systemic therapy were reviewed. As resection of metastases is a controversial therapeutic approach, no clinical trials are available for review. Rather, evidence for a potential role for surgery rests on retrospective studies of small series of patients. Technical advances have rendered resection of liver and lung metastases safe. Long-term results as reported by other investigators support the role of metastasectomy in selected patients. The site of failure following ablation of liver metastases is usually in the liver. Following resection of lung metastases, nonpulmonary and disseminated recurrences are most common. Adjuvant therapy with docetaxel or any other agent or combination with significant activity against visceral metastases might potentiate long-term results.

L12 ANSWER 54 OF 55 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
ACCESSION NUMBER: 2001:77815 BIOSIS

DOCUMENT NUMBER: PREV200100077815
TITLE: A phase II trial of escalated dose docetaxel (TXT) with G-CSF support in patients (pts) with advanced breast cancer.
AUTHOR(S): Mitchell, P. [Reprint author]; Bassar, R.; Harris, M. [Reprint author]; Ng, S.; Gibbs, P. [Reprint author]; Chipman, M. [Reprint author]; Grigg, A.; Jeffrey, A.; James, R.; Gargano, J.; Riva, A.; Appia, F.; Green, M.
CORPORATE SOURCE: Medical Oncology, Austin and Repatriation Medical Centre, Heidelberg West, VIC, Australia
SOURCE: Breast Cancer Research and Treatment, (November, 2000) Vol. 64, No. 1, pp. 88. print.
Meeting Info.: 23rd Annual San Antonio Breast Cancer Symposium. San Antonio, Texas, USA. December 06-09, 2000. Cancer Therapy and Research Center Research Foundation.
CODEN: BCTR6. ISSN: 0167-6806.
DOCUMENT TYPE: Conference; (Meeting)
Conference; Abstract; (Meeting Abstract)
LANGUAGE: English
ENTRY DATE: Entered STN: 7 Feb 2001
Last Updated on STN: 12 Feb 2002

L12 ANSWER 55 OF 55 BIOSIS COPYRIGHT (c) 2006 The Thomson Corporation on STN
ACCESSION NUMBER: 2001:132253 BIOSIS
DOCUMENT NUMBER: PREV200100132253
TITLE: Close correlation of paraneoplastic hyperfibrinolysis with relapse and remission of anaplastic small cell carcinoma: A case report.
AUTHOR(S): Kegel, T. [Reprint author]; Kellner, O. [Reprint author]; Grothey, A. [Reprint author]; Wolf, H.-H. [Reprint author]; Voigt, W. [Reprint author]; Dorligshaw, O. [Reprint author]; Schmoll, H.-J. [Reprint author]
CORPORATE SOURCE: Dept. of Hematology/Oncology, University of Halle, Halle, Germany
SOURCE: Onkologie, (October, 2000) Vol. 23, No. Sonderheft 7, pp. 184. print.
Meeting Info.: Annual Meeting of the German and Austrian Society for Hematology and Oncology. Graz, Austria. October 21-25, 2000.
CODEN: ONKOD2. ISSN: 0378-584X.
DOCUMENT TYPE: Conference; (Meeting)
Conference; Abstract; (Meeting Abstract)
LANGUAGE: English
ENTRY DATE: Entered STN: 14 Mar 2001
Last Updated on STN: 15 Feb 2002